



Iris
Biotech

...using either our standard reagent or a modified biotinylation agent. It is also possible to incorporate biotin at a specific position. Our selection of reagents for reversible biotinylation includes a **biotinylation reagent** for the required synthesis, a selection of **Biotin-derivatives** and options if you are looking for a special derivative not listed, or for a custom synthesis.

Biotinylation - In some cases, reversible biotinylation can be cleaved using reducing agents such as **biotin-binding proteins** (e.g. streptavidin) which can bind to biotin.

BIOTINYLLATION REAGENTS



Version: IB2_4

Empowering Peptide Innovation

With this guiding theme in mind, Iris Biotech's mission is to support researchers by supplying

- innovative technologies,
- rare compounds,
- as well as a broad portfolio on standard consumables,

available in flexible quantities from small scale to bulk quantities. To fulfill our dedication "Empowering Peptide Innovation", we are attending various conferences, symposia, and exhibitions each year. This allows us to remain in direct contact with scientists all over the world, both from academia and industry, to exchange knowledge, and to gather new ideas to tackle your current challenges.

Guided by our dedication to provide

- competent service,
- as well as novel substances and
- latest technologies,

Iris Biotech is your trusted partner for the world of peptides, while having strong expertise in associated disciplines. Thus, our portfolio comprises reagents and tools for the synthesis and modification of peptides, e.g., amino acids, resins and solvents but also for related technologies such as drug delivery, linkerology® and life sciences.

Owed to the growing demand for tailor-made compounds, our portfolio is fine-tuned by our custom synthesis service at Iris Biotech Laboratories. Our skilled scientists offer profound expertise in

- *de novo* route development,
- upscaling towards larger scale production,
- as well as synthesis optimization for increased efficiency.

Examples are the synthesis of rare chiral building blocks, unnatural amino acid derivatives, sophisticated orthogonal protecting groups, heterocycles, building blocks for nucleotides, PEGs and PEG-analogs as well as specific linkers for controlled drug delivery and release.



Amino Acids



Building Blocks



Life Sciences



Drug Delivery



Reagents



Resins



Linkerology®



Click Chemistry

Portfolio Overview

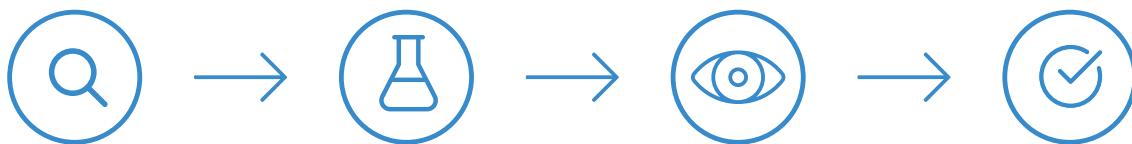
Peptide Synthesis and Modification	Linkerology® and Drug Delivery	Life Sciences
(Protected) Amino Acids Standards such as Fmoc-D/L-AAA and Boc-D/L-AAA, Smoc amino acids for peptide synthesis in water, variety of protecting groups (e.g., Pbf, Trt, ^t Bu, Bzl, Acm, Mob, SIT, Phacm, Allocam, Mmt), unusual amino acids, fluorinated derivatives, substituted prolines, arginine analogs	Linkers for Solid Phase Peptide Synthesis Cleavable Linkers Val-Ala-based, Val-Cit-based, disulfide-based, Dde-helping hands, pH-sensitive linkers	Biotinylation Reagents Carbohydrates Galactose, Glucose, Mannose, Xylose and others
Building Blocks Amino alcohols, amino aldehydes, diamines and hydrazines, (pseudoproline) dipeptides, polyamines and spermines, fatty acid derivatives, peptide nucleic acids (PNAs)	Photo-Activatable Linkers Functionalized Linkers Clickable linkers, trifunctional linkers, linkers with maleimide function, cross-linkers, selective N-term acylation and biotinylation, 5HP2O	Drug Metabolites Peptides Substrates & Inhibitors E.g., protein kinase inhibitors, substrates for fusion (Halo/Snap/Clip)-tagged proteins
Reagents Coupling reagents, solvents and scavengers, protecting groups	PROTACs Ligands, linkers & modules	Natural Products Dyes and Fluorescent Labels E.g., ICG, AMC, DAPI
Resins Preloaded resins (e.g., based on Trityl, TCP, TentaGel, Methoxybenzhydryl, Merrifield, PAM, Rink, Wang), scavenger resins, hydrazone resins, poly(acrylamide) resins, Cyclover	Fullerenes, Poly(2-oxazolines), Dextrans & Plant-Derived Cholesterol Superparamagnetic Iron Oxide Nanoparticles Poly-Amino Acids Poly-Arg, Poly-Glu, Poly-Lys, Poly-Orn, Poly-Sar	Maillard & Amadori Reaction Products Large portfolio of derivatives useful as standards for food, pharma and cosmetics industry
	PEGylation Branched PEGylating reagents, (amino-)PEG-acids, PEG-amines & hydrazides & guanidines, reagents for Click-conjugation, Biotin-PEG-reagents, PEG-thiols, PEG-maleimides, other PEGylating reagents	Vitamins

Custom Synthesis

Your project requires a compound not listed in our portfolio?
Get in contact and inquire about our custom synthesis capabilities.

Our experienced scientists are excited to accept your synthetic challenge!

In such cases, your request undergoes the following stages:



Step-by-Step Analysis Process Evaluation

- Customer's demands
- Detailed literature review
- Synthetic possibilities

Strategy Development Quality Consistency

- Protocol development
- Method development and validation
- Customized synthesis
- Identity confirmation
- Purity verification

Our Service Promise

All our services are based on high standards, transparency & documentation, trust, honesty & confidentiality, as well as the required know-how.

High Standards

- Values: sustainability & responsibility
- State-of-the-art equipment & latest technologies
- High quality standards
- Qualified suppliers & regular audits

Transparency & Documentation

- Talk to our specialists – customer care
- Certificates of analysis & origin
- Impurity profiling
- Safety data sheets
- Analytical and process reports

Trust, Honesty & Confidentiality

- Intergenerational business valuing partnerships
- Meeting the customer's expectations
- Integrity towards our customers

Our Know-How

- One-step reactions & complex multi-step synthesis
- Scalability from mg to kg quantities
- Route scouting



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Biotinylation Reagents

1. Background

Biotinylation describes the attachment of biotin to a protein or other macromolecule. The selective isolation of such biotin-containing molecules out of complex mixtures is based on the high affinity of avidin and streptavidin for biotin, representing one of the strongest known non-covalent interactions between protein and ligand ($K_d = 10^{-15}$ M) and being unaffected by variations of e.g., pH, temperature, or organic solvents. Furthermore, biotin is a rather small molecule compared to other labels, minimizing its steric impact on the biotinylated carrier itself. Thus, this nonradioactive method represents a highly important and versatile tool for purification, immobilization, and labeling.

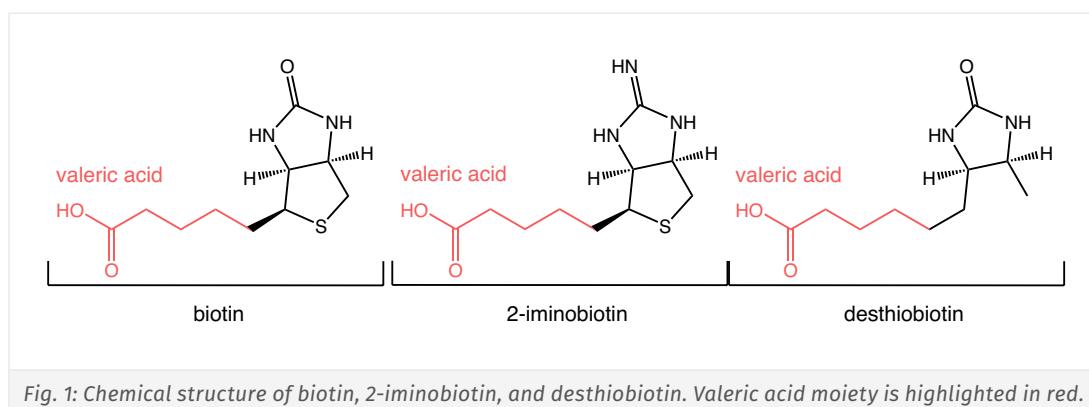


Fig. 1: Chemical structure of biotin, 2-iminobiotin, and desthiobiotin. Valeric acid moiety is highlighted in red.

As biotin bears a valeric acid side-chain, it can easily be derivatized without affecting its (strept)avidin-binding site. The reactivity can be tailored towards various functional groups, e.g., primary amines, sulphydryls and carboxyls, as well as derivatives with various linker lengths to adjust the availability of biotin for (strept)avidin binding, with incorporated PEG moieties for improved solubility, with cleavable conjugates, or for reversible biotin labelling. This range of possibilities allows to perfectly suit the biotinylation reagent for the required application.

2-Iminobiotin and desthiobiotin also bind tightly to (strept)avidin, however, not as strong as biotin. Compounds labelled with 2-iminobiotin and desthiobiotin can therefore be gently stripped from accordingly labeled targets with buffered biotin solutions avoiding harsh denaturation conditions such as 6 M GdmCl.



**Iris
Biotech**



You need further details about our
custom synthesis capabilities?

Get in contact!

📞 +49 (0) 9231 97121-0
📠 +49 (0) 9231 97121-99
✉️ info@iris-biotech.de
🌐 www.iris-biotech.de

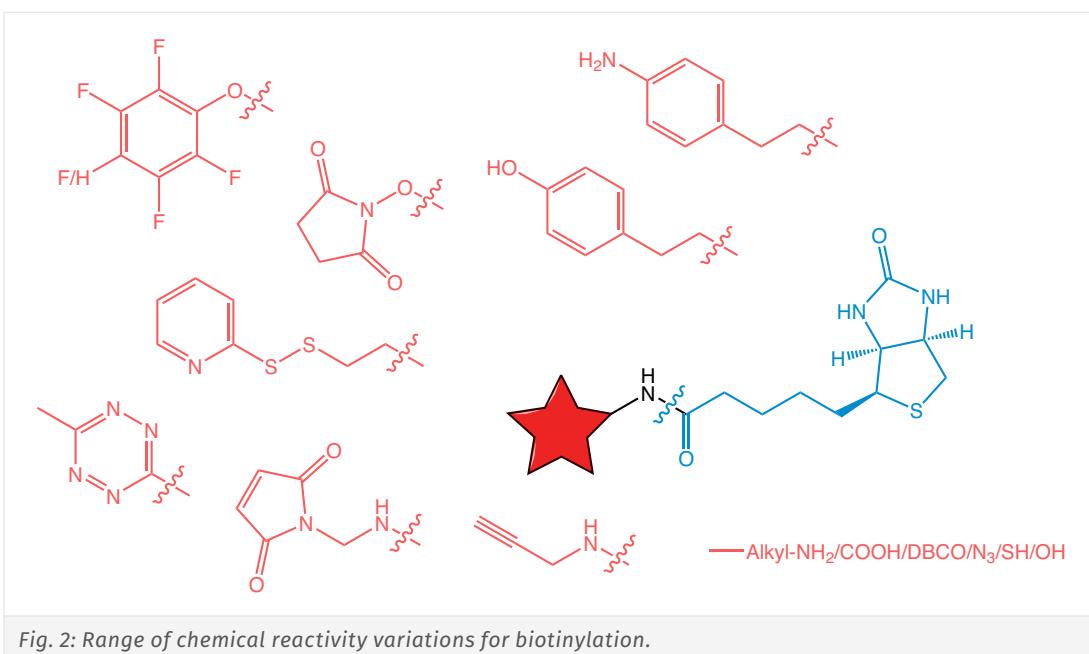


Fig. 2: Range of chemical reactivity variations for biotinylation.

Selecting the right biotinylation reagent to use for a specific application and protein of interest depends on a number of factors that must be carefully considered and optimized, including:

- Solubility – Modifications in biotinylation reagent solubility allow access to target proteins in hydrophobic or hydrophilic environments and influence the solubility of the biotinylated protein.
- Spacer arm length – The availability of biotin for (strept)avidin binding might be influenced by the length of the spacer arm.
- Cleavability/reversibility – Captured biotinylated proteins can be recovered or purified by cleaving the biotin molecule from the target protein or reversing attenuated biotin-(strept)avidin interactions.
- Functional group – Specific reactive moieties bind to certain amino acids or to the functional groups of specific amino acids for nonselective or targeted biotinylation, respectively.

Overcoming the Hydrophobic Nature of Biotin

Typical biotinylation with biotin-NHS normally results in multiple attachments, as several lysines are usually present on protein surfaces. Hence, the hydrophobic properties of biotin will influence surface polarity and solubility in aqueous conditions. In case such biotin labelling has a negative impact on the above listed protein properties, incorporation of a PEG unit between the conjugation point and biotin itself is a standard method of choice to maintain or improve physical properties of the modified protein.

PEGs are molecules comprised of repeating ethylene glycol units with solubility enhancing properties both in hydrophobic and hydrophilic environment. PEGs are considered as non-immunogenic and show promising results in numerous applications. They have a dramatic impact on protein solubility, stability, and aggregation, and improve ELISAs, IHC assays, and formulations in many aspects.

Background

Amine-Reactive
Biotinylation Reagents

Carboxy-Reactive
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Reagents for Various Protein
and RNA Biotinylations

Reversible Biotinylation

Sulfhydryl-Reactive
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Biotinylation Reagents

Some protein-biotin conjugates are particularly susceptible to aggregation and loss of antigen binding ability, which can complicate assays and provide irreproducible and/or erroneous results. In comparison, biotinylated IgGs employing a PEG linker show much lower tendency for aggregation, maintain a high affinity for the antigen, and the resulting complexes with the biotin-binding proteins remain completely soluble.

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2. Amine-Reactive Biotinylation Reagents

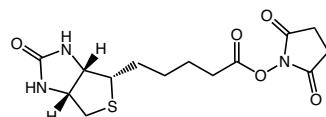
Active esters such as N-hydroxysuccinimide (NHS) esters permit a catalyst-free conjugation reaction to amines, e.g., of lysine side-chains and the N-terminus, under mild conditions. Yet, NHS esters may hydrolyze over time if, for example, atmospheric moisture has condensed on the substance. As alternatives, tetrafluorophenyl (TFP) and pentafluorophenyl (PFP) active esters can be applied that are much more resistant to hydrolysis.

[Product details](#)

RL-1006 Biotin-OSu

N-Hydroxysuccinimidobiotin

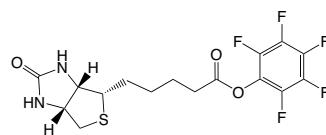
CAS-No.	35013-72-0
Formula	C ₁₄ H ₁₉ N ₃ O ₅ S
Mol. weight	341,39 g/mol



RL-8615 Biotin-PFP

Biotin pentafluorophenyl ester

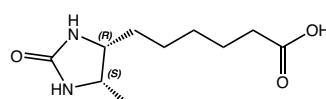
CAS-No.	120550-35-8
Formula	C ₁₆ H ₁₅ F ₅ N ₂ O ₃ S
Mol. weight	410,36 g/mol



LS-1650 D-Desthiobiotin

(4R,5S)-5-methyl-2-oxo-4-Imidazolidinehexanoic acid

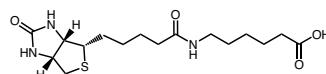
CAS-No.	533-48-2
Formula	C ₁₀ H ₁₈ N ₂ O ₃
Mol. weight	214,27 g/mol



RL-2025 Biotin-Ahx-OH

6-Biotinylamino-hexanoic acid

CAS-No.	72040-64-3
Formula	C ₁₆ H ₂₇ N ₃ O ₄ S
Mol. weight	357,47 g/mol


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Biotinylation Reagents

Product details

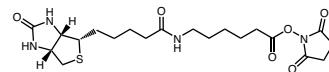
RL-2020 Biotin-Ahx-NHS

6-Biotinylamino-hexanoic acid-N-hydroxysuccinimidyl ester

CAS-No. 72040-63-2

Formula C₂₀H₃₀N₄O₆S

Mol. weight 454,54 g/mol



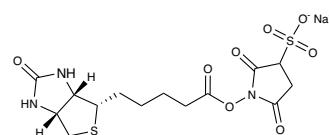
LS-1680 Sulfo-NHS-Biotin

Biotin 3-sulfo-N-hydroxysuccinimide ester sodium salt

CAS-No. 194041-65-1

Formula C₁₄H₁₈N₃NaO₈S₂

Mol. weight 443,42 g/mol



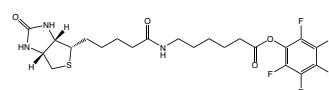
RL-3880 Biotin-Aca-PFP

Biotin-6-aminohexanoic acid pentafluorophenyl ester

CAS-No. 273207-85-5

Formula C₂₂H₂₆F₅N₃O₄S

Mol. weight 523,52 g/mol



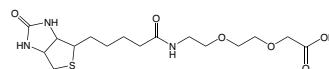
PEG2560 Biotin-O₂Oc-OH

8-Biotinylamido-3,6-dioxaoctanoic acid

CAS-No. 1238575-77-3

Formula C₁₈H₂₇N₃O₆S

Mol. weight 389,47 g/mol



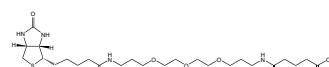
PEG5280 Biotin-TOTA-glutaramic acid*DIPEA

5,21-dioxo-25-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)-10,13,16-trioxa-6,20-diaza-pentacosan-1-oic acid diisopropylethylamine salt

CAS-No. 1205744-09-7

Formula C₂₅H₄₄N₄O₈S*C₈H₁₉N

Mol. weight 560,70*129,30 g/mol



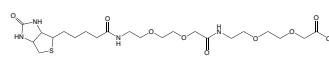
PEG2565 Biotin-O₂Oc-O₂Oc-OH

8-Biotinylamido-3,6-dioxaoctanoic acid dimer

CAS-No. 1301706-65-9

Formula C₂₂H₃₈N₄O₈S

Mol. weight 534,62 g/mol



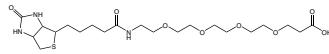
**PEG1515 Biotin-PEG(4)-COOH**

15-Biotinamino-4,7,10,13-tetraoxa-pentadecanoic acid

CAS-No. 721431-18-1

Formula C₂₁H₃₇N₃O₈S

Mol. weight 491,6 g/mol

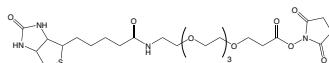
**PEG1870 Biotin-PEG(4)-NHS**

15-Biotinamino-4,7,10,13-tetraoxa-pentadecanoic acid succinimidyl ester

CAS-No. 459426-22-3

Formula C₂₅H₄₀N₄O₁₀S

Mol. weight 588,67 g/mol

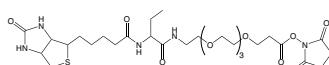
**PEG1845 Biotin-PEG(4)-NHS-(Biotinidase resistant)**

18-Biotinamino-17-oxo-4,7,10,13-tetraoxa-16-azaicosan-1-oic acid succinimidyl ester

CAS-No. 1334172-61-0

Formula C₂₉H₄₇N₅O₁₁S

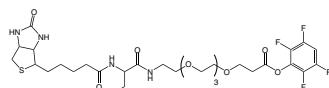
Mol. weight 673,78 g/mol

**PEG7760 Biotin-PEG(4)-TFP-(Biotinidase resistant)**

18-Biotinamino-17-oxo-4,7,10,13-tetraoxa-16-azaicosan-1-oic acid (2,3,5,6-tetrafluorophenyl) ester

Formula C₃₁H₄₄F₄N₄O₉S

Mol. weight 724,76 g/mol

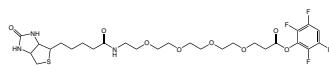
**PEG2071 Biotin-TEG-TFP**

Biotin-tetra(ethylene glycol)-2,3,5,6-tetrafluorophenyl ester

CAS-No. 2985664-57-9

Formula C₂₇H₃₇F₄N₃O₈S

Mol. weight 639,66 g/mol

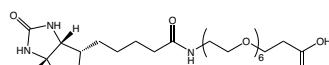
**PEG6690 Biotin-PEG(6)-COOH**

21-[D(+)-Biotinylamino]-4,7,10,13,16,19-hexaoxaheneicosanoic acid

CAS-No. 1352814-10-8

Formula C₂₅H₄₅N₃O₁₀S

Mol. weight 579,7 g/mol



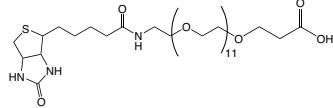
Biotinylation Reagents

Product details

PEG1051 Biotin-PEG(12)-COOH

alpha-Biotin-omega-(propionic acid)-dodeca(ethylene glycol)

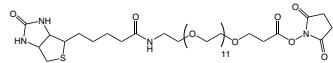
CAS-No. 948595-11-7
Formula C₃₇H₆₉N₃O₁₆S
Mol. weight 844,04 g/mol



PEG1860 Biotin-PEG(12)-NHS

alpha-Biotin-omega-carboxy succinimidyl ester dodeca(ethylene glycol)

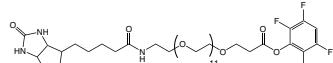
CAS-No. 365441-71-0
Formula C₄₁H₇₂N₄O₁₈S
Mol. weight 941,09 g/mol



PEG5130 Biotin-PEG(12)-TFP

alpha-Biotin-omega-propionic acid (2,3,5,6-tetrafluorophenyl) ester dodeca(ethylene glycol)

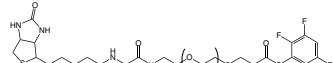
CAS-No. 3017961-69-9
Formula C₄₃H₆₉F₄N₃O₁₆S
Mol. weight 992,08 g/mol



PEG7770 Biotin-PEG(12)-TFP-(Biotinidase resistant)

alpha-Biotinyl-(2-aminobutyryl)-omega-(2,3,5,6-tetrafluorophenyl propionate) dodeca(ethylene glycol)

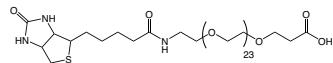
Formula C₄₇H₇₆F₄N₄O₁₇S
Mol. weight 1077,18 g/mol



PEG4260 Biotin-PEG(24)-COOH

alpha-Biotin-omega-(propionic acid) 24(ethylene glycol)

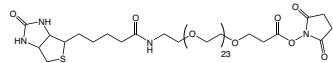
CAS-No. 721431-18-1
Formula C₆₁H₁₁₇N₃O₂₈S
Mol. weight 1372,65 g/mol



PEG4250 Biotin-PEG(24)-NHS

alpha-Biotin-omega-(succinimidyl propionate) 24(ethylene glycol)

CAS-No. 365441-71-0
Formula C₆₅H₁₂₀N₄O₃₀S
Mol. weight 1469,72 g/mol

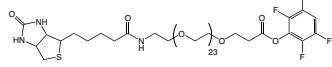


Product details

PEG7750 Biotin-PEG(24)-TFP

alpha-Biotin-omega-(2,3,5,6-tetrafluorophenyl propionate) 24(ethylene glycol)

Formula C₆₇H₁₁₇F₄N₃O₂₈S
Mol. weight 1520,71 g/mol


PEG4270 Biotin-PEG(48)-COOH

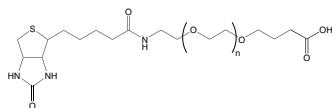
alpha-Biotin-omega-(propionic acid) 48(ethylene glycol)

CAS-No. 721431-18-1
Formula C₁₁₂H₂₁₈N₄O₅₃S
Mol. weight 2500,99 g/mol


PEG1053 Biotin-PEG-COOH (3 kDa)

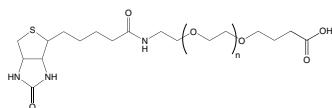
alpha-Biotin-omega-carboxy poly(ethylene glycol)

Mol. weight 3000 Da


PEG1054 Biotin-PEG-COOH (5 kDa)

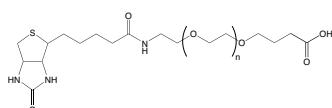
alpha-Biotin-omega-carboxy poly(ethylene glycol)

Mol. weight 5000 Da


PEG1052 Biotin-PEG-COOH (10 kDa)

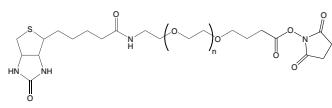
alpha-Biotin-omega-carboxy poly(ethylene glycol)

Mol. weight 10000 Da


PEG1056 Biotin-PEG-NHS (3 kDa)

alpha-Biotin-omega-carboxy succinimidyl ester poly(ethylene glycol)

Mol. weight 3000 Da



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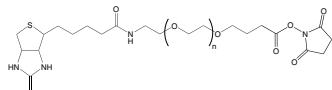
Biotinylation Reagents

Product details

PEG1057 Biotin-PEG-NHS (5 kDa)

alpha-Biotin-omega-carboxy succinimidyl ester poly(ethylene glycol)

Mol. weight 5000 Da



PEG1055 Biotin-PEG-NHS (10 kDa)

alpha-Biotin-omega-carboxy succinimidyl ester poly(ethylene glycol)

Mol. weight 10000 Da

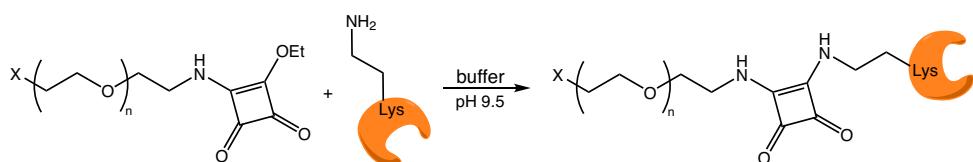
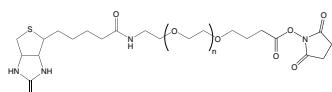


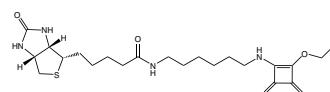
Fig. 3: Squaric acid ethyl esters chemoselectively bind to the amino side-chain functions of lysine in proteins, whereas the guanidine function of arginine, the hydroxyl groups of serine and threonine and the imidazole function of histidine are not affected.

Product details

RL-8430 Biotin-Hx-SQA

N-(6-((2-ethoxy-3,4-dioxocyclobut-1-en-1-yl)amino)hexyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide

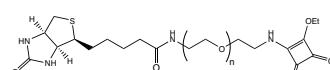
Formula C₂₂H₃₄N₄O₅S
Mol. weight 466,60 g/mol



PEG6590 Biotin-PEG-SQA (3 kDa)

alpha-Biotin-omega-squaric acid ethyl ester poly(ethylene glycol)

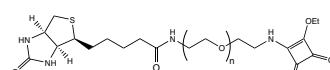
Mol. weight 3000 Da



PEG6595 Biotin-PEG-SQA (5 kDa)

alpha-Biotin-omega-squaric acid ethyl ester poly(ethylene glycol)

Mol. weight 5000 Da

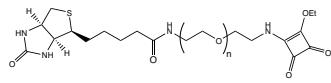


Product details

PEG6580 Biotin-PEG-SQA (10 kDa)

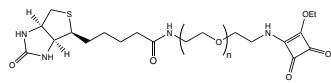
alpha-Biotin-omega-squaric acid ethyl ester poly(ethylene glycol)

Mol. weight 10000 Da

**PEG6585 Biotin-PEG-SQA (20 kDa)**

alpha-Biotin-omega-squaric acid ethyl ester poly(ethylene glycol)

Mol. weight 20000 Da

**References:**

- *Synthesis and reactivity of compounds with cyclobutane ring(s). 10. Syntheses of squaric acid, its monoorthesters, and related derivatives via [2 + 2] cycloadditions of tetraalkoxyethylenes with heterosubstituted ketenes; D. Bellus; J. Org. Chem. 2002; **44**: 1208-1211.* ↗ <https://doi.org/10.1021/jo01322a004>
- *Be squared: expanding the horizon of squaric acid-mediated conjugations; F. R. Wurm, H. A. Klok; Chem Soc Rev 2013; **42**: 8220-36.* ↗ <https://doi.org/10.1039/c3cs60153f>

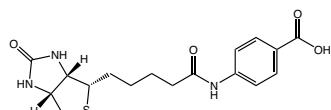
Biotin-PABA is a carboxylic acid terminated biotin derivative and as such amine reactive for chemical biotinylation of amino functions, e.g., of lysines. Its main use is as a substrate for biotinidase, which cleaves biotin-amide to release biotin *in vivo*. As byproduct, *para*-amino benzoic acid is released, which can be quantified by either fluorescent or colorimetric methods.

Product details

LS-4200 Biotin-PABA

4-(5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamido)benzoic acid

CAS-No. 6929-40-4
 Formula C₁₇H₂₁N₃O₄S
 Mol. weight 363,43 g/mol

**Reference:**

- *Qualitative colorimetric ultramicroassay for the detection of biotinidase deficiency in newborns; E. C. Gonzalez, N. Marrero, A. Frometa, D. Herrera, E. Castells, P. L. Perez; Clin Chim Acta 2006; **369**: 35-9.*
 ↗ <https://doi.org/10.1016/j.cca.2006.01.009>

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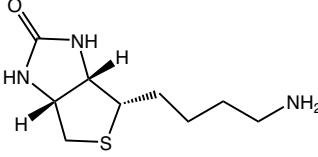
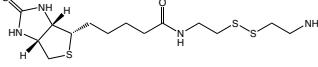
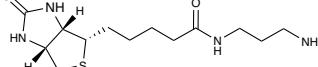
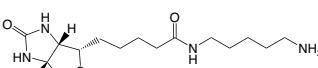
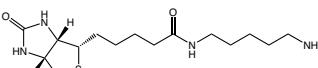
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3. Carboxy-Reactive Biotinylation Reagents

Carboxy groups are present on aspartic acid and glutamate side-chains, as well as on the carboxy-terminal ends, and can be reacted with amino-bearing biotinylation reagents in the presence of a coupling reagent, e.g., EDC.

		Product details
RL-8435	Norbiotinamine*HCl	(3aS,4S,6aR)-4-(4-aminobutyl)tetrahydro-1H-thieno[3,4-d]imidazol-2(3H)-one hydrochloride
CAS-No.	160385-86-4	
Formula	C ₉ H ₁₇ N ₃ OS*HCl	
Mol. weight	215,32*36,45 g/mol	
RL-3391	Biotin-cystamine*TFA	Biotinamidoethylidisulfanyl(2-aminoethane) trifluoroacetate
CAS-No.	880491-09-8	
Formula	C ₁₄ H ₂₆ N ₄ O ₂ S ₃ *CF ₃ COOH	
Mol. weight	378,58*114,02 g/mol	
LS-1690	Biotinyl-DAPr*TFA	N-(+)-Biotinyl-3-aminopropylammonium trifluoroacetate
CAS-No.	244760-26-7	
Formula	C ₁₃ H ₂₄ N ₄ O ₂ S*CF ₃ CO ₂ H	
Mol. weight	300,42*114,02 g/mol	
RL-2030	Biotin cadaverine nTFA	5-(Biotinamido)pentylamine nTFA
CAS-No.	288259-39-2	
Formula	C ₁₅ H ₂₈ N ₄ O ₂ S nC ₂ HF ₃ O ₂	
Mol. weight	328,47 net g/mol	
LS-1670	Biotinyl cadaverine	N-(5-aminopentyl)biotinamide
CAS-No.	115416-38-1	
Formula	C ₁₅ H ₂₈ N ₄ O ₂ S	
Mol. weight	328,48 g/mol	

Product details

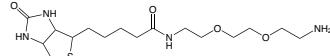
PEG2760 Biotin-DOOA*HCl

1-Biotinyl-3,6-dioxa-8-octaneamine hydrochloride

CAS-No. 862373-14-6

Formula $C_{16}H_{30}N_4O_4S^*HCl$

Mol. weight 374,50*36,45 g/mol

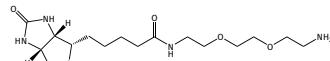

RL-4060 Biotin-DOOA

Biotinyl-1-amino-3,6-dioxa-8-octanamine

CAS-No. 138529-46-1

Formula $C_{16}H_{30}N_4O_4S$

Mol. weight 374,50 g/mol

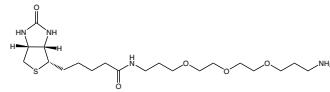

PEG8050 Biotin-TOTA

N-(13-amino-4,7,10-trioxatridecanyl)-D-biotinamide

CAS-No. 183896-00-6

Formula $C_{20}H_{38}N_4O_5S$

Mol. weight 446,6 g/mol

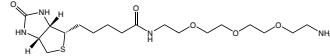

RL-4210 Biotin-PEG(3)-NH₂

N-[2-[2-[2-(2-Aminoethoxy)ethoxy]ethoxy]ethyl]biotinamide

CAS-No. 359860-27-8

Formula $C_{18}H_{34}N_4O_5S$

Mol. weight 418,22 g/mol

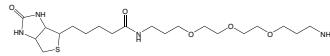

PEG2110 Biotin-PEG(3)-NH₂*TFA

N-Biotin-3,3'-(2,2'-oxybis(ethane-2,1-diyl)bis(oxo))dipropan-1-amine trifluoroacetate

CAS-No. 1334172-59-6

Formula $C_{20}H_{38}N_4O_5S^*CF_3CO_2H$

Mol. weight 446,60*114,01 g/mol

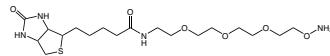

PEG5140 Biotin-TEG-O-NH₂*HCl

N-(2-(2-(2-(aminoxy)ethoxy)ethoxy)ethoxy)ethyl-5-biotinamide hydrochloride

CAS-No. 1951424-88-6

Formula $C_{18}H_{34}N_4O_6S^*HCl$

Mol. weight 434,55*36,45 g/mol



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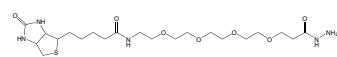
PEG1425 Biotin-PEG(4)-NHNH₂

15-Biotinamino-4,7,10,13-tetraoxa-pentadecanoyl hydrazide

CAS-No. 756525-97-0

Formula C₂₁H₃₉N₅O₇S

Mol. weight 505,63 g/mol



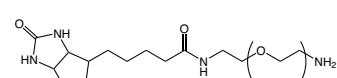
PEG4290 Biotin-PEG(7)-NH₂

alpha-Biotin-omega-amino hepta(ethylene glycol)

CAS-No. 1334172-76-7

Formula C₂₆H₅₀N₄O₉S

Mol. weight 594,76 g/mol



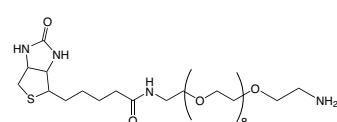
PEG1044 Biotin-PEG(9)-NH₂

alpha-Biotin-omega-amino-nona(ethylene glycol)

CAS-No. 960132-48-3

Formula C₃₀H₅₈N₄O₁₁S

Mol. weight 682,88 g/mol



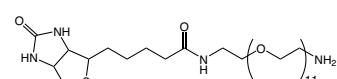
PEG4300 Biotin-PEG(11)-NH₂

alpha-Biotin-omega-amino undeca(ethylene glycol)

CAS-No. 604786-74-5

Formula C₃₄H₆₆N₄O₁₃S

Mol. weight 770,97 g/mol

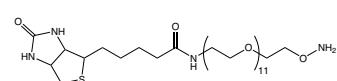


PEG4680 Biotin-PEG(11)-O-NH₂*HCl

alpha-Biotinyl-omega-oxyamine-undeca(ethylene glycol) hydrochloride

Formula C₃₄H₆₆N₄O₁₄S

Mol. weight 786,97*36,45 g/mol



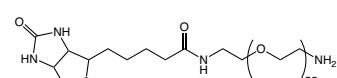
PEG4310 Biotin-PEG(23)-NH₂

alpha-Biotin-omega-amino 23(ethylene glycol)

CAS-No. 604786-74-5

Formula C₅₈H₁₁₄N₄O₂₅S

Mol. weight 1299,6 g/mol



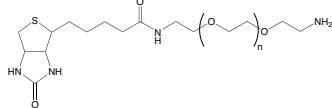
Product details

PEG1046 Biotin-PEG-NH₂ (3 kDa)

alpha-Biotin-omega-amino poly(ethylene glycol)

CAS-No. 604786-74-5

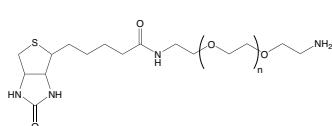
Mol. weight 3000 Da

**PEG1047 Biotin-PEG-NH₂ (5 kDa)**

alpha-Biotin-omega-amino poly(ethylene glycol)

CAS-No. 604786-74-5

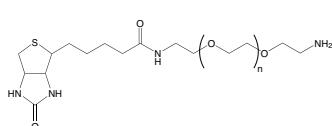
Mol. weight 5000 Da

**PEG1045 Biotin-PEG-NH₂ (10 kDa)**

alpha-Biotin-omega-amino poly(ethylene glycol)

CAS-No. 604786-74-5

Mol. weight 10000 Da



Find more information and background on PEGylation in our booklet PEGylation!


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4. Reagents for Various Protein and RNA Biotinylations

Tyramine Derivatives

Tyramine derivatives such as biotin-tyramide are converted to radicals by horseradish peroxidase (HRP) in the presence of H₂O₂. These highly reactive species preferentially react with tyrosine residues of proteins, a property that is employed in many applications such as protein labeling, *in situ* hybridization, electron microscopy, ELISA, phage display, and others. Biotin-tyramide is a valuable tool for signal amplification and can significantly improve signal-to-noise ratios.

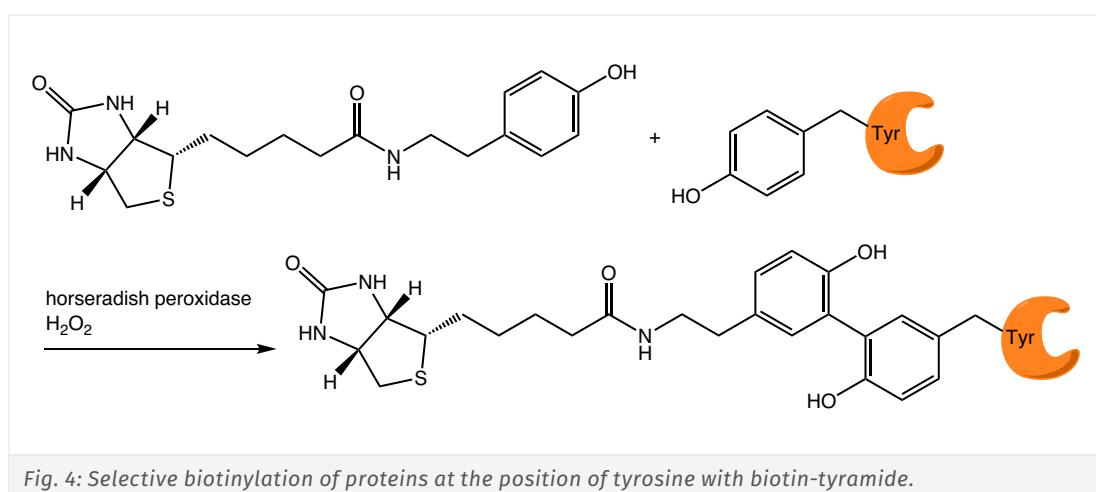
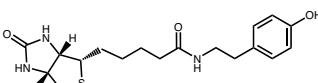
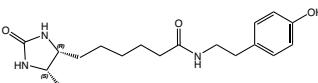
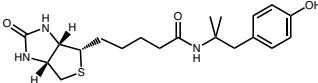
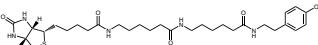
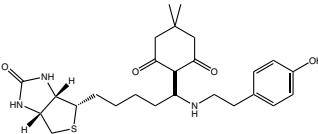


Fig. 4: Selective biotinylation of proteins at the position of tyrosine with biotin-tyramide.

		Product details
LS-3500	Biotin Tyramide	 N-(3aS,4S,6aR)-hexahydro-N-[2-(4-hydroxyphenyl)ethyl]-2-oxo-1H-thieno[3,4-d]imidazole-4-pentanamide CAS-No. 41994-02-9 Formula C ₁₈ H ₂₅ N ₃ O ₃ S Mol. weight 363,47 g/mol
LS-1660	Desthiobiotin-Tyramide	 N-(4-hydroxyphenethyl)-6-((4R,5S)-5-methyl-2-oxoimidazolidin-4-yl)hexanamide CAS-No. 2242902-55-0 Formula C ₁₈ H ₂₇ N ₃ O ₃ Mol. weight 333,43 g/mol

		Product details
LS-4060	Biotin-Dimethyl-Tyramide	 N-(1-(4-hydroxyphenyl)-2-methylpropan-2-yl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide CAS-No. 2808072-95-7 Formula C ₂₀ H ₂₉ N ₃ O ₃ S Mol. weight 391,53 g/mol
LS-4030	Biotin-Ahx-Ahx-Tyramide	 Biotin-Ahx-Ahx-Tyramide CAS-No. 851113-28-5 Formula C ₃₀ H ₄₇ N ₅ O ₃ S Mol. weight 589,8 g/mol
LS-4000	Biotin-Dde-Tyramide	 2-(1-(4-hydroxyphenethylamino)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentylidene)-5,5-dimethylcyclohexane-1,3-dione CAS-No. 2819732-80-2 Formula C ₂₆ H ₃₅ N ₃ O ₄ S Mol. weight 485,64 g/mol
PEG8130	Biotin-PEG(4)-Dde-Tyramide	 N-(15-(4,4-dimethyl-2,6-dioxocyclohexylidene)-18-(4-hydroxyphenyl)-3,6,9,12-tetraoxa-16-azaoctadecyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide CAS-No. 2814457-48-0 Formula C ₃₄ H ₅₆ N ₄ O ₉ S Mol. weight 732,93 g/mol

References:

- New insights into the DT40 B cell receptor cluster using a proteomic proximity labeling assay; X. W. Li, J. S. Rees, P. Xue, H. Zhang, S. W. Hamaia, B. Sanderson, P. E. Funk, R. W. Farndale, K. S. Lilley, S. Perrett, A. P. Jackson; *J Biol Chem* 2014; **289**: 14434-47. ↗ <https://doi.org/10.1074/jbc.M113.529578>
- Tyramide signal amplification for analysis of kinase activity by intracellular flow cytometry; M. R. Clutter, G. C. Heffner, P. O. Krutzik, K. L. Sachen, G. P. Nolan; *Cytometry A* 2010; **77**: 1020-31.
↗ <https://doi.org/10.1002/cyto.a.20970>
- Proteomic mapping of mitochondria in living cells via spatially restricted enzymatic tagging; H. W. Rhee, P. Zou, N. D. Udeshi, J. D. Martell, V. K. Mootha, S. A. Carr, A. Y. Ting; *Science* 2013; **339**: 1328-1331.
↗ <https://doi.org/10.1126/science.1230593>
- A catalysis-based selection for peroxidase antibodies with increased activity; J. Yin, J. H. Mills, P. G. Schultz; *J Am Chem Soc* 2004; **126**: 3006-7. ↗ <https://doi.org/10.1021/ja0391980>
- Catalyzed reporter deposition, a novel method of signal amplification. II. Application to membrane immunoassays; M. N. Bobrow, K. J. Shaughnessy, G. J. Litt; *J Immunol Methods* 1991; **137**: 103-12.
↗ [https://doi.org/10.1016/0022-1759\(91\)90399-z](https://doi.org/10.1016/0022-1759(91)90399-z)

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Biotinylation Reagents

- Catalyzed reporter deposition, a novel method of signal amplification. Application to immunoassays; M. N. Bobrow, T. D. Harris, K. J. Shaughnessy, G. J. Litt; *J Immunol Methods* 1989; **125**: 279-85.
↗ [https://doi.org/10.1016/0022-1759\(89\)90104-x](https://doi.org/10.1016/0022-1759(89)90104-x)
- The Oxidation of Tyramine, Tyrosine, and Related Compounds by Peroxidase; A. J. Gross, I. W. Sizer; *J. Biol. Chem.* 1959; **234**: 1611-1614.
- WO2008128352 A1.

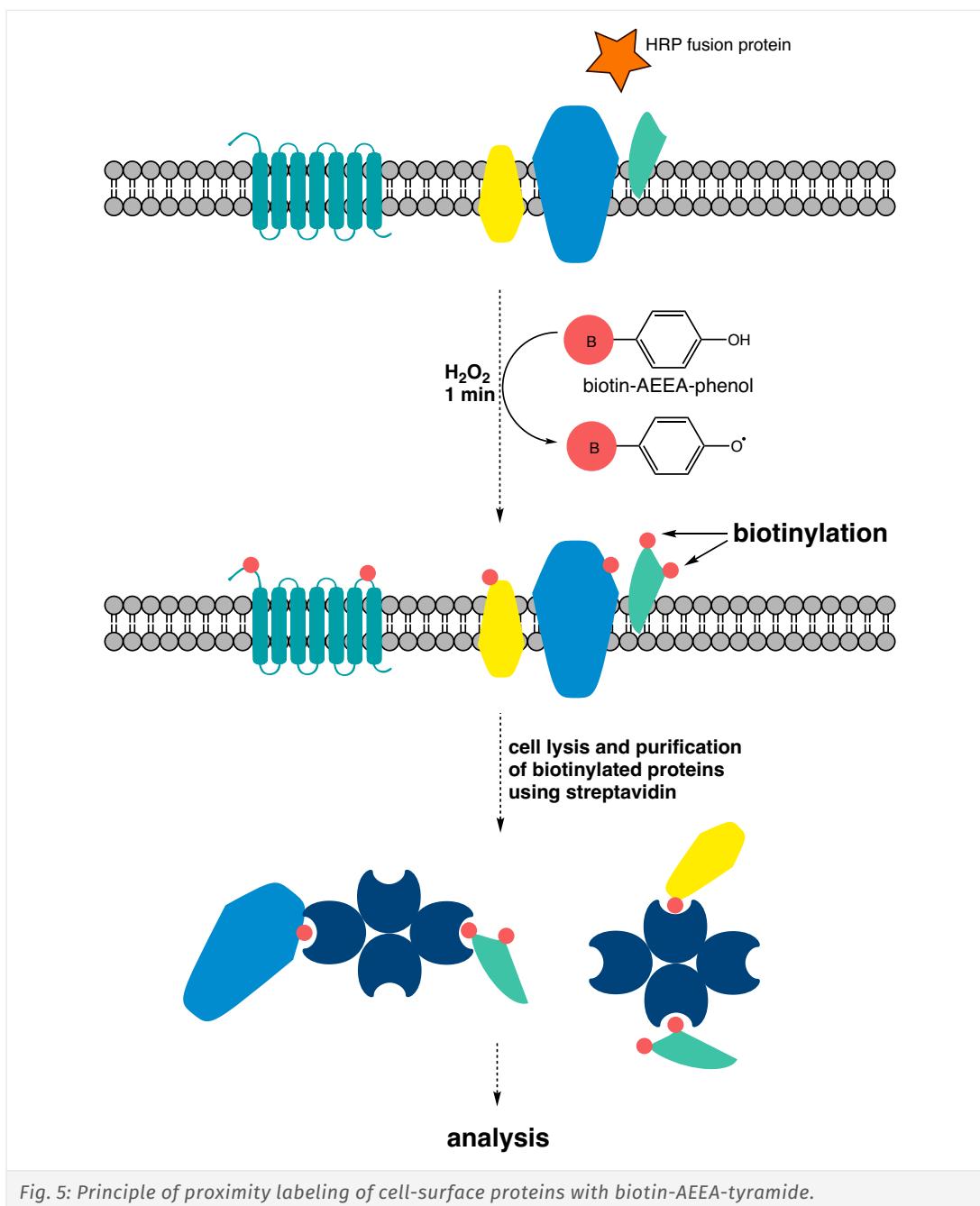


Fig. 5: Principle of proximity labeling of cell-surface proteins with biotin-AEEA-tyramide.

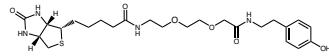
Hydrophilic biotin-AEEA-tyramide, which bears a short PEG spacer between the biotin and the phenol moieties, is shown to be a membrane-impermeant variant of regular biotin-tyramide when used for a short time on cultured cells (ca. 1 min during the biotinylation reaction). Biotin-AEEA-tyramide can thus be used for the selective biotinylation of cell surface proteins.

Product details

LS-3490 Biotin-AEEA-Tyramide

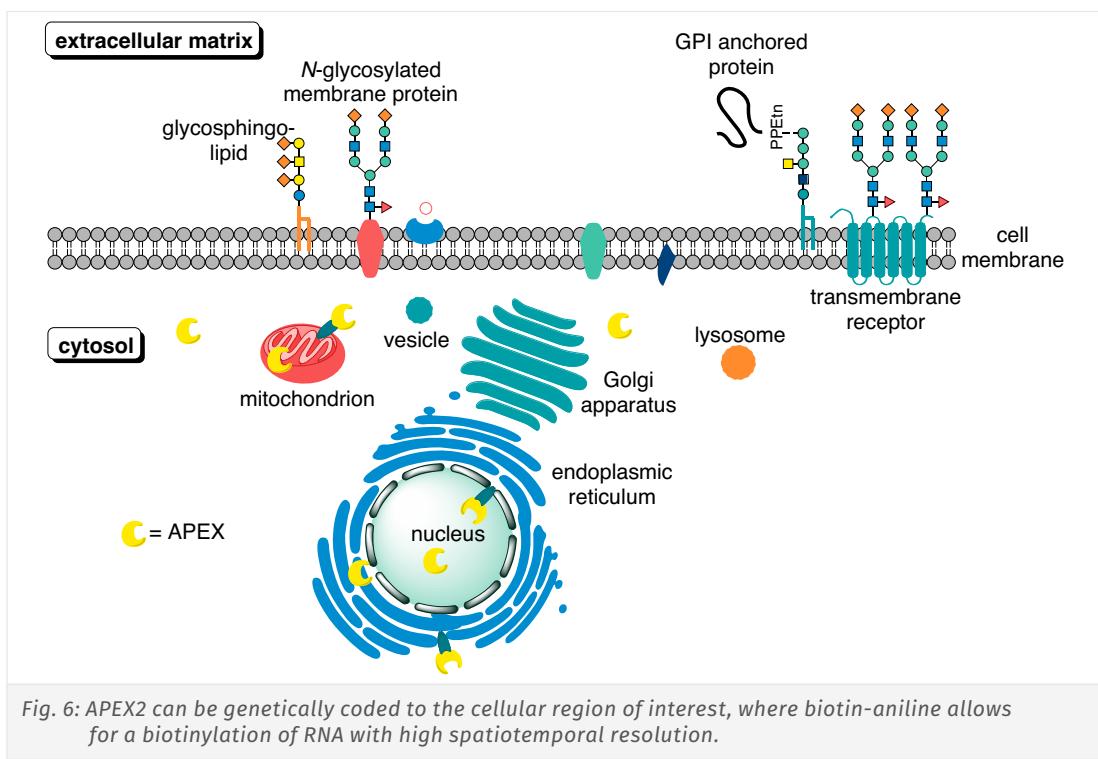
N-(2-(2-(4-hydroxyphenethylamino)-2-oxethoxy)ethoxyethyl)-5((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide

CAS-No. 2561473-92-3
 Formula C₂₄H₃₆N₄O₆S
 Mol. weight 508,63 g/mol


Reference:

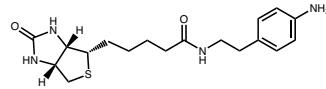
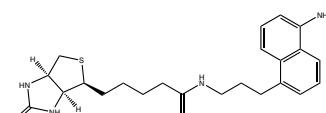
- *Mapping the Proteome of the Synaptic Cleft through Proximity Labeling Reveals New Cleft Proteins; T. Cijssouw, A. Ramsey, T. Lam, B. Carbone, T. Blanpied, T. Biederer; Proteomes 2018; 6: 48.*
 <https://doi.org/10.3390/proteomes6040048>

Proximity labeling of proteins with biotin-phenol (biotin-tyramide) and an engineered peroxidase enzyme (e.g., APEX = Engineered Ascorbate Peroxidase) is a pivotal tool for molecular biology. So far, mainly phenolic compounds have been used as substrates for APEX enzymes. The enzymatic reaction yields phenoxyl radicals that predominantly react with electron-rich amino acid side-chains. Consequently, phenolic compounds conjugated to biotin are not ideal for the biotinylation of nucleic acids. In order to facilitate the efficient labeling of nucleic acids, arylamine-biotin conjugates were developed (Zhou et al., Angew. Chem. Int. Ed. 2019). These novel probes show a significantly higher reactivity towards nucleic acids compared to biotin-phenol. In combination with the peroxidase APEX2, these probes allow for the mapping of the subcellular transcriptome by proximity-dependent labeling in living cells, and may aid in elucidating the subcellular organization of biomolecules. When combined with RNA sequencing (APEX-seq), this labeling method provides complete sequence information for various classes of RNA with a high spatiotemporal resolution (Fazal & Han et al., Cell 2019).



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Biotinylation Reagents

		Product details
LS-3970	Biotin-aniline N-(4-aminophenethyl)-biotinamide CAS-No. 769933-15-5 Formula C ₁₈ H ₂₆ N ₄ O ₂ S Mol. weight 362,49 g/mol	 
LS-4650	Biotin-naphthylamine N-(3-(5-aminonaphthalen-1-yl)propyl)-D-biotinamide CAS-No. 2375201-60-6 Formula C ₂₃ H ₃₀ N ₄ O ₂ S Mol. weight 426,58 g/mol	 

References:

- Expanding APEX2 Substrates for Proximity-Dependent Labeling of Nucleic Acids and Proteins in Living Cells; Y. Zhou, G. Wang, P. Wang, Z. Li, T. Yue, J. Wang, P. Zou; *Angew Chem Int Ed* 2019; **58**: 11763-11767. <https://doi.org/10.1002/anie.201905949>
- Atlas of Subcellular RNA Localization Revealed by APEX-Seq; F. M. Fazal, S. Han, K. R. Parker, P. Kaewsapsak, J. Xu, A. N. Boettiger, H. Y. Chang, A. Y. Ting; *Cell* 2019; **178**: 473-490 e26. <https://doi.org/10.1016/j.cell.2019.05.027>
- Directed evolution of APEX2 for electron microscopy and proximity labeling; S. S. Lam, J. D. Martell, K. J. Kamer, T. J. Deerinck, M. H. Ellisman, V. K. Mootha, A. Y. Ting; *Nat Methods* 2015; **12**: 51-4. <https://doi.org/10.1038/nmeth.3179>

SNAP-tag® and CLIP-tag™

Biotin-conjugated benzylguanine (BG) reacts specifically and rapidly with the so-called SNAP-tag®, a polypeptide protein tag, allowing irreversible and covalent labeling of SNAP fusion proteins with biotin. Biotinylated proteins can for example be selectively isolated based on the high affinity towards (strept) avidin representing a useful tool for purification, immobilization, and labeling.

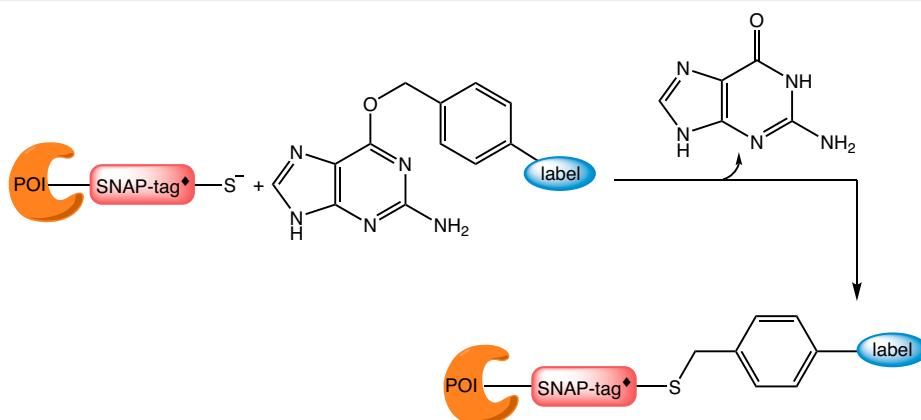
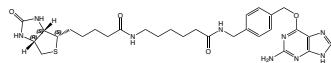


Fig. 7: The SNAP-tag® is a 20 kDa self-labeling protein tag based on a modified form of the human O6-alkylguanine-DNA-alkyltransferase (hAGT), a DNA repair enzyme. A cysteine residue within the SNAP-tag® undergoes an irreversible reaction with synthetic O6-benzylguanine (BG) derivatives resulting in a covalent thioether bond. The BG moiety can easily be further functionalized with a label of choice, e.g., fluorophore, biotin, generally without affecting the reaction of the substrate with the SNAP-tag®.

Product details

RL-3860 Biotin-SNAP

N-(4-(((2-amino-9H-purin-6-yl)oxy)methyl)benzyl)-6-(5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamido)hexanamide



CAS-No. 471918-16-8
 Formula C₂₉H₃₉N₉O₄S
 Mol. weight 609,75 g/mol


References:

- *SNAP-Tag Technology: A Useful Tool To Determine Affinity Constants and Other Functional Parameters of Novel Antibody Fragments*; J. Niesen, M. Sack, M. Seidel, R. Fendel, S. Barth, R. Fischer, C. Stein; *Bioconjug Chem* 2016; **27**: 1931-41. ↗ <https://doi.org/10.1021/acs.bioconjchem.6b00315>
- *Site-specific, covalent labeling of recombinant antibody fragments via fusion to an engineered version of 6-O-alkylguanine DNA alkyltransferase*; F. Kampmeier, M. Ribbert, T. Nachreiner, S. Dembski, F. Beaufils, A. Brecht, S. Barth; *Bioconjug Chem* 2009; **20**: 1010-5. ↗ <https://doi.org/10.1021/bc9000257>
- *A general method for the covalent labeling of fusion proteins with small molecules in vivo*; A. Keppler, S. Gendreizig, T. Gronemeyer, H. Pick, H. Vogel, K. Johnsson; *Nat Biotechnol* 2003; **21**: 86-9.
 ↗ <https://doi.org/10.1038/nbt765>
- *Directed Evolution of O6-Alkylguanine-DNA Alkyltransferase for Efficient Labeling of Fusion Proteins with Small Molecules In Vivo*; A. Juillerat, T. Gronemeyer, A. Keppler, S. Gendreizig, H. Pick, H. Vogel, K. Johnsson; *Chem. Biol.* 2003; **10**: 313-317. ↗ [https://doi.org/10.1016/s1074-5521\(03\)00068-1](https://doi.org/10.1016/s1074-5521(03)00068-1)

Biotin-conjugated benzylcytosine (BC) reacts specifically and rapidly with the so-called CLIP-tag™, a polypeptide protein tag, allowing irreversible and covalent labeling of clip fusion proteins with biotin. Biotinylated proteins can for example be selectively isolated based on the high affinity towards (strept) avidin representing a useful tool for purification, immobilization, and labeling.

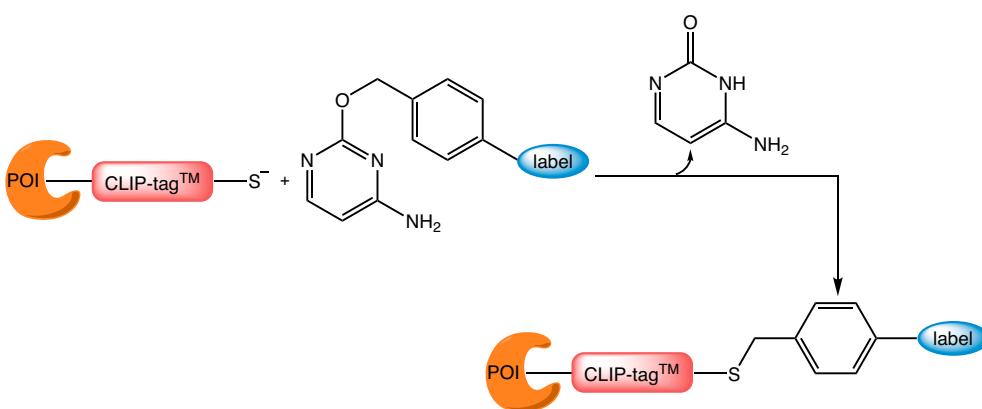


Fig. 8: The CLIP-tag™ (20 kDa) is a modified version of the SNAP-tag®, engineered to react with benzylcytosine (BC) instead of benzylguanine (BG). Thus, properties are similar. CLIP-tag™- and SNAP-tag®-fused proteins can be labeled simultaneously in the same cells.

SNAP-tag® is a registered trademark and CLIP-tag™ a trademark of New England Biolabs.
 His Tag® is a registered trademark of Merck KGAA.

Background

Amine-Reactive Biotinylation Reagents

Carboxy-Reactive Biotinylation Reagents

Reagents for Various Protein and RNA Biotinylations

Reversible Biotinylation

Sulfhydryl-Reactive Biotinylation Reagents

Click-Reactive Biotinylation Reagents

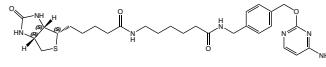
Biotinylated Amino Acids

Other Biotin Reagents

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Biotinylation Reagents

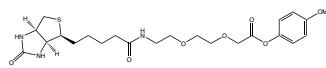
		Product details
RL-3870	Biotin-Clip N-(4-(((4-aminopyrimidin-2-yl)oxy)methyl)benzyl)-6-(5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamido)hexanamide  CAS-No. 1004524-73-5 Formula C ₂₈ H ₃₉ N ₇ O ₄ S Mol. weight 569,73 g/mol	

Reference:

- A general method for the covalent labeling of fusion proteins with small molecules *in vivo*; A. Keppler, S. Gendreizig, T. Gronemeyer, H. Pick, H. Vogel, K. Johnsson; **Nat Biotechnol** 2003; **21**: 86-9.
 <https://doi.org/10.1038/nbt765>

His-tag

This 4-methoxyphenyl ester adds selectively to Gly-His-tags of proteins, while it does not react with amino functions of surface accessible lysines. Therefore, it expresses a unique possibility to label proteins regiospecifically at the Gly-His-tag position, enabling a wide application field in chemical biology and for biopharmaceuticals.

		Product details
RL-3100	Biotin-AEEA-OPhOMe 2-(2-(Biotinamido)ethoxy)ethoxyacetic acid 4-methoxyphenyl ester  CAS-No. 2546513-67-9 Formula C ₂₃ H ₃₃ N ₃ O ₅ S Mol. weight 495,59 g/mol	

For more details about the methodology for highly selective N-terminal chemical acylation of expressed proteins, please look at our blog at: www.iris-biotech.de/blog/potm-his-tags-for-the-chemical-modification-of-peptides/

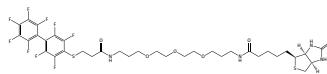
References:

- Selective N-terminal acylation of peptides and proteins with a Gly-His tag sequence; M. C. Martos-Maldonado, C. T. Hjuler, K. K. Sorensen, M. B. Thygesen, J. E. Rasmussen, K. Villadsen, S. R. Midtgård, S. Kol, S. Schöffelen, K. J. Jensen; **Nat Commun** 2018; **9**: 3307.  <https://doi.org/10.1038/s41467-018-05695-3>
- Selective Acylation of Proteins at Gly and Lys in His Tags; K. J. Jensen, M. B. Thygesen, K. K. Sørensen, S. Wu, T. Treiberg, S. Schöffelen; **ChemBioChem** 2022; **23(24)**: e202200359.  <https://doi.org/10.1002/cbic.202200359>

Another way for site-selective conjugation is the so-called π-Clamp-mediated cysteine conjugation. Pentelute and co-workers showed that the incorporation of the four-amino-acid sequence Phe-Cys-Pro-Phe, the so-called „π-Clamp“, tunes the reactivity of this cysteine thiol for site-selective conjugation with perfluoroaromatic reagents. Thus, this technology can be used to selectively modify one cysteine site in proteins containing multiple endogenous cysteine residues.

RL-4050 PFB-mercaptopropionyl-TOTA-Biotin

Perfluorobiphenyl-mercaptopropionyl-TOTA-Biotin

 Formula C35H41F9N4O6S2
 Mol. weight 848,84 g/mol
**References:**

- Enzymatic „click“ ligation: selective cysteine modification in polypeptides enabled by promiscuous glutathione S-transferase; C. Zhang, A. M. Spokoyny, Y. Zou, M. D. Simon, B. L. Pentelute; *Angew Chem Int Ed Engl.* 2013; **52**(52): 14001-5. <https://doi.org/10.1002/anie.201306430>
- Convergent diversity-oriented side-chain macrocyclization scan for unprotected polypeptides; Y. Zou, A. M. Spokoyny, C. Zhang, M. D. Simon, H. Yu, Y.-S. Lin, B. L. Pentelute; *Org. Biomol. Chem.* 2014; **12**: 566-573. <https://doi.org/10.1039/C3OB42168F>
- π-Clamp-mediated cysteine conjugation; C. Zhang, M. Welborn, T. Zhu, N. J. Yang, M. S. Santos, T. Van Voorhis, B. L. Pentelute; *Nat Chem.* 2016; **8**: 120-128. <https://doi.org/10.1038/nchem.2413>
- Site-Specific Small Molecule Labeling of an Internal Loop in JC Polyomavirus Pentamers Using the π-Clamp-Mediated Cysteine Conjugation; J. A. Baccile, P. J. Voorhees, A. J. Chillo, M. Berry, R. Morgenstern, T. J. Schwertfeger, F. M. Rossi, C. D. S. Nelson; *ChemBioChem* 2022; **(21)**: 3037-3041. <https://doi.org/10.1002/cbic.202100188>

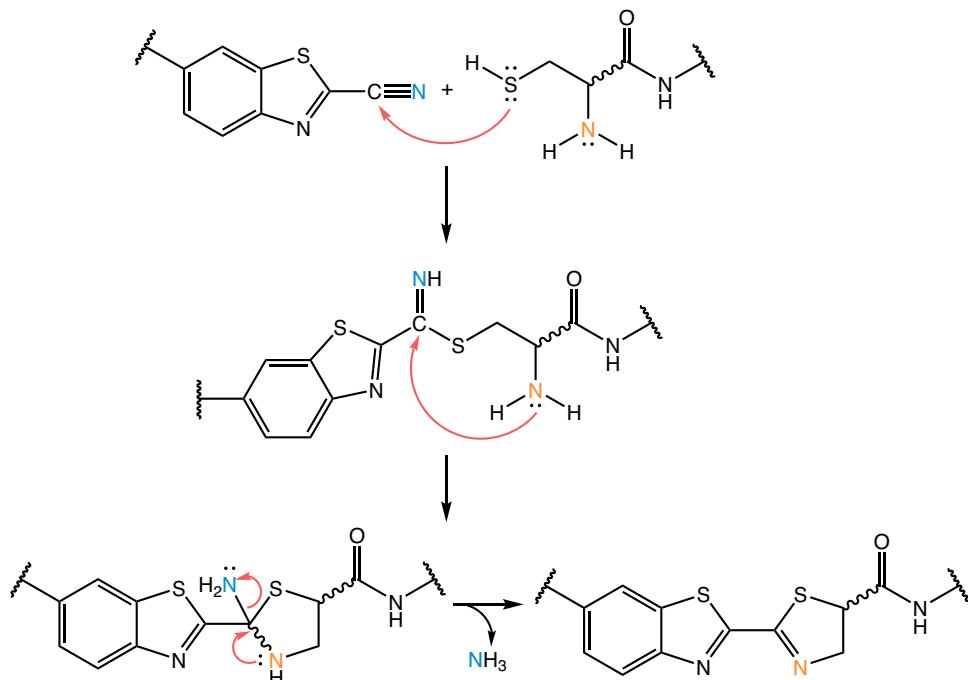
CBT Derivatives

Fig. 9: Mechanism of the CBT-Click reaction: The electron pair of the sulfur attacks the carbon of the cyano group, the nitrogen forms an enamine. The cysteine's nitrogen then attacks the carbon of the enamine, and a cyclic intermediate is formed. Finally, the lone electron pair of the amino nitrogen of the cysteine attacks the positively charged carbon to yield the product. The nitrogen which came from the nitrile is leaving the reaction as ammonia.

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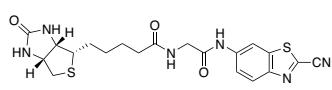
Biotinylation Reagents

Product details

RL-4230 Biotin-Gly-CBT

N-(2-((2-cyanobenzo[d]thiazol-6-yl)amino)-2-oxoethyl)-biotin

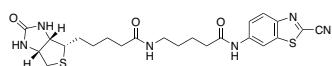
Formula C₂₀H₂₂N₆O₃S₂
Mol. weight 458,56 g/mol



RL-4240 Biotin-Pen-CBT

N-(5-((2-cyano-6-benzothiazolyl)amino)-5-oxopentyl)-biotin

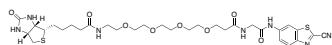
CAS-No. 2550397-17-4
Formula C₂₃H₂₆N₆O₃S₂
Mol. weight 500,64 g/mol



RL-4250 Biotin-PEG(4)-Gly-CBT

15-Biotinylamino-4,7,10,13-tetraoxa-pentadecanoyl-glycyl-N-2-cyano-benzothiazole

CAS-No. 1907639-46-6
Formula C₃₁H₄₃N₇O₈S₂
Mol. weight 705,85 g/mol



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5. Reversible Biotinylation

Competitive Displacement of Desthiobiotin or 2-Iminobiotin by Biotin

The single-ring, sulfur-free desthiobiotin binds to (strept)avidin with almost the same specificity as biotin but has a lower affinity than biotin ($K_d=10^{-11}$ M vs. $K_d=10^{-15}$ M, respectively). Thus, milder elution conditions based on competitive displacement with free biotin are sufficient for the efficient release of desthiobiotinylated proteins from (strept)avidin. In addition, as further advantage, endogenous biotinylated molecules remain bound to (strept)avidin during pull-down assay experiments with biological samples, which allows easy differentiation and prevents co-purification.

In comparison to biotin and desthiobiotin, the guanidino analog of biotin (= 2-iminobiotin) shows an even weaker binding affinity for (strept)avidin, which is, in addition, pH-dependent. At high pH (> 9), iminobiotin-tagged proteins bind to (strept)avidin conjugates with high affinity, but the (strept)avidin-iminobiotin complexes dissociate at low pH (~ 4) or via replacement upon addition of biotin. This allows the purification of a captured protein without denaturation, releasing biologically functional, 2-iminobiotinylated-proteins from the (strept)avidin conjugates. Besides, 2-iminobiotin reversibly inhibits nitric oxide synthases (NOS). NOS oxidizes the guanidino-nitrogen of L-arginine, resulting in L-citrulline and nitric oxide.

Tab.1: Overview of different biotin derivatives and affinity towards (strept)avidin.

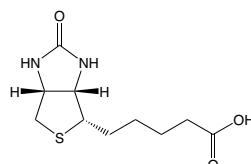
Compound	Biotin	Desthiobiotin	2-Iminobiotin
Structure			
Affinity towards (strept)avidin (K_d)	10^{-15} M	10^{-11} M	10^{-8} M

Product details

LS-1070 D-Biotin

Hexahydro-2-oxo-1H-thieno[3,4-d]imidazole-4-pentanoic acid

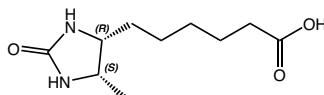
CAS-No. 58-85-5
Formula $C_{10}H_{16}N_2O_3S$
Mol. weight 244,31 g/mol



LS-1650 D-Desthiobiotin

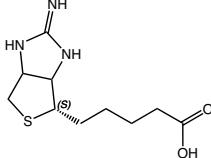
(4R,5S)-5-methyl-2-oxo-4-imidazolidinehexanoic acid

CAS-No. 533-48-2
Formula $C_{10}H_{16}N_2O_3$
Mol. weight 214,27 g/mol



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Biotinylation Reagents

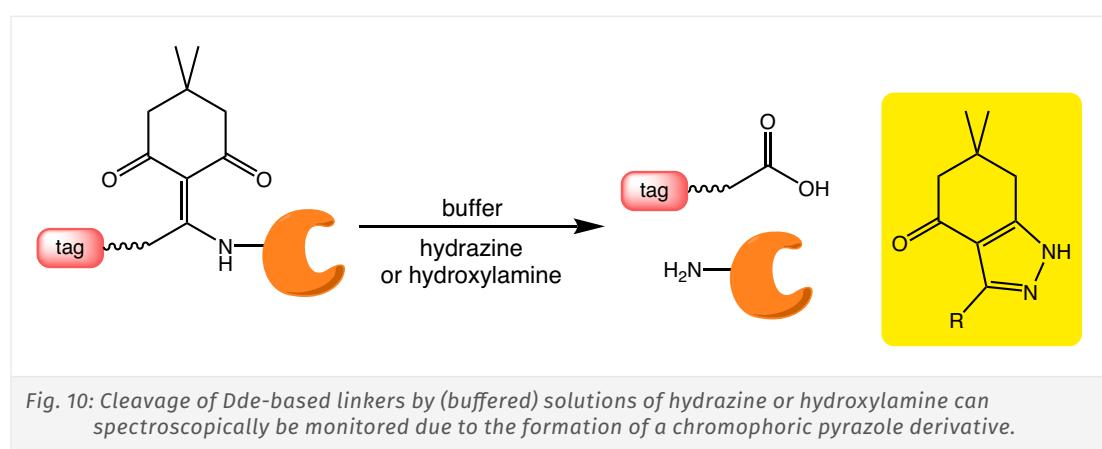
		Product details
LS-4320	2-Iminobiotin	<p>5-((4S)-2-iminohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanoic acid</p> <p>CAS-No. 13395-35-2 Formula C₁₀H₁₇N₃O₂S Mol. weight 243,32 g/mol</p>  

References:

- Iminobiotin affinity columns and their application to retrieval of streptavidin; K. Hofmann, S. W. Wood, C. C. Brinton, J. A. Montibeller, F. M. Finn; *Proc Natl Acad Sci U S A* 1980; **77**: 4666-8.
 <https://doi.org/10.1073/pnas.77.8.4666>
- Voltammetric homogeneous binding assay of biotin without a separation step using iminobiotin labeled with an electroactive compound; K. Sugawara, N. Kamiya, G. Hirabayashi, H. Kuramitz; *Anal. Sci.* 2005; **21**: 897-900.  <https://doi.org/10.2116/analsci.21.897>
- Easily reversible desthiobiotin binding to streptavidin, avidin, and other biotin-binding proteins: uses for protein labeling, detection, and isolation; J. D. Hirsch, L. Eslamizar, B. J. Filanowski, N. Malekzadeh, R. P. Haugland, J. M. Beechem, R. P. Haugland; *Anal Biochem* 2002; **308**: 343-57.
 [https://doi.org/10.1016/s0003-2697\(02\)00201-4](https://doi.org/10.1016/s0003-2697(02)00201-4)
- The use of the 2-iminobiotin-avidin interaction for the selective retrieval of labeled plasma membrane components; G. A. Orr; *J. Biol. Chem.* 1981; **256**: 761-766.  [https://doi.org/10.1016/s0021-9258\(19\)70041-6](https://doi.org/10.1016/s0021-9258(19)70041-6)

Dde-based Linkers

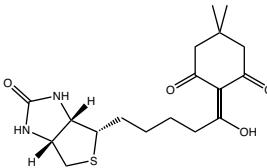
Placing Dde as one terminal group of a linker and a functional group prone for conjugation as the other, or using Dde as the central connective portion of a linker, allows for the creation of new bifunctional and cleavable linkers. Dde can be selectively cleaved by using a buffered, aqueous solution of hydrazine or hydroxylamine, thus cleaving the linkage after (strept)avidin attachment. The cleavage can be monitored spectroscopically, as the resulting pyrazole shows a strong absorption at 290 nm.



LS-4020 Biotin-Dde

2-(1-hydroxy-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentylidene)-5,5-dimethylcyclohexane-1,3-dione

CAS-No. 194038-08-9
 Formula C₁₈H₂₆N₂O₄S
 Mol. weight 366,48 g/mol



Background

Amine-Reactive
Biotinylation ReagentsCarboxy-Reactive
Biotinylation ReagentsReagents for Various Protein
and RNA Biotinylations

Reversible Biotinylation

Sulfhydryl-Reactive
Biotinylation ReagentsClick-Reactive
Biotinylation Reagents

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Dde-based linkers can selectively and temporarily be attached to:

- biomolecules for binding to (strept)avidin (with terminal biotin) (A);
- conjugation to any solid supports, e.g., via Click reaction (B);
- solubilizing tags, e.g., hexa-lysine (C) or oligo-arginine;
- PEGs (D) or other hydrophilic groups improving the solubility;
- dyes and any other conjugate for monitoring, diagnostics, targeting or other purposes.

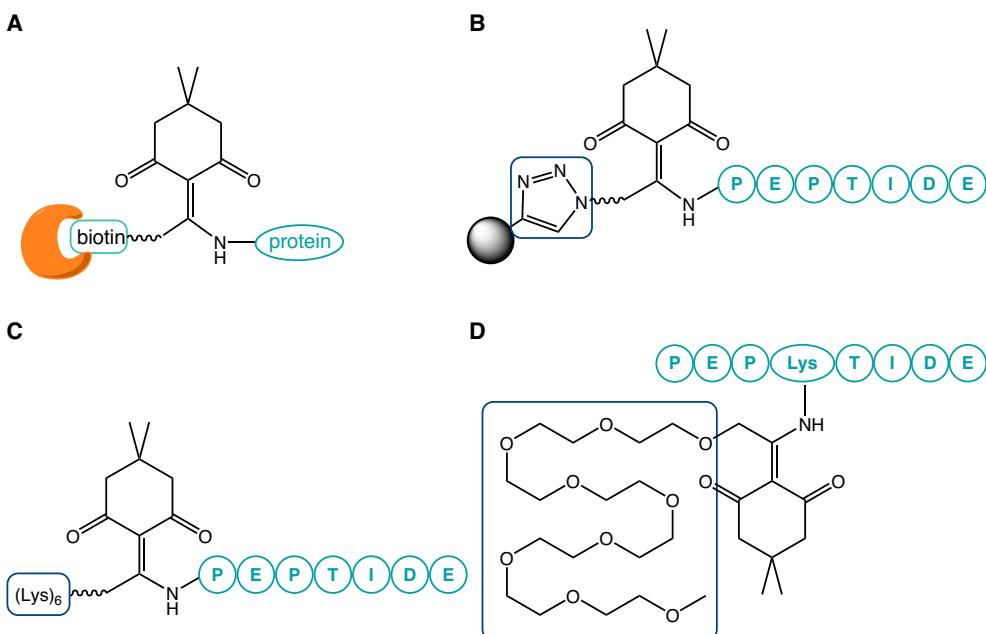


Fig. 11: Dde-based linkers can be utilized for various applications: Attachment of a cleavable biotin tag to proteins for catch-and-release affinity purification over (strept)avidin beads (A), reversible labeling or conjugation to other biomolecules, or reversible immobilization on solid supports via Click chemistry (B), temporary attachment of solubilizing tags like oligo-lysine (C) or PEGs (D).

Dde-based linkers are one option of cleavable linkers with mild cleavage conditions. A different widely applicable methodology is the introduction of disulfides. Biotinyling variants with cleavable linkers based on disulfides can be cleaved using reducing agents such as glutathione. Alternatively, reducing conditions inside certain cell compartments enable specific elution of biotinylated molecules from biotin-binding proteins.

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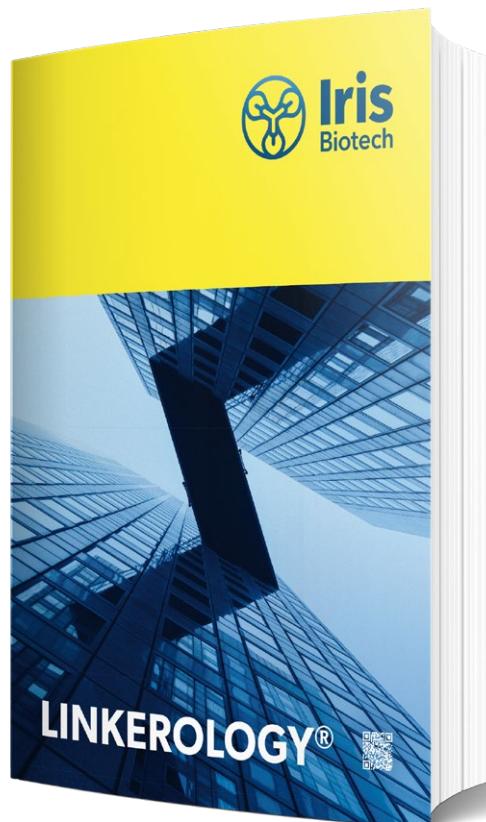
Biotinylation Reagents

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- Synthesis of hydrophobic insulin-based peptides using a helping hand strategy; M. M. Disotuar, M. E. Petersen, J. M. Nogueira, M. S. Kay, D. H. Chou; *Org Biomol Chem* 2019; **17**: 1703-1708. ↗ <https://doi.org/10.1039/c8ob01212a>
- Mapping the Binding Site of BMS-708163 on gamma-Secretase with Cleavable Photoprobes; N. Gertsik, C. W. Am Ende, K. F. Geoghegan, C. Nguyen, P. Mukherjee, S. Mente, U. Seneviratne, D. S. Johnson, Y. M. Li; *Cell Chem Biol* 2017; **24**: 3-8. ↗ <https://doi.org/10.1016/j.chembiol.2016.12.006>
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Biotin-SS-NHS is a disulfide bridge-containing biotinylation construct that can be cleaved using reducing agents such as glutathione or by reducing conditions inside certain cell compartments.

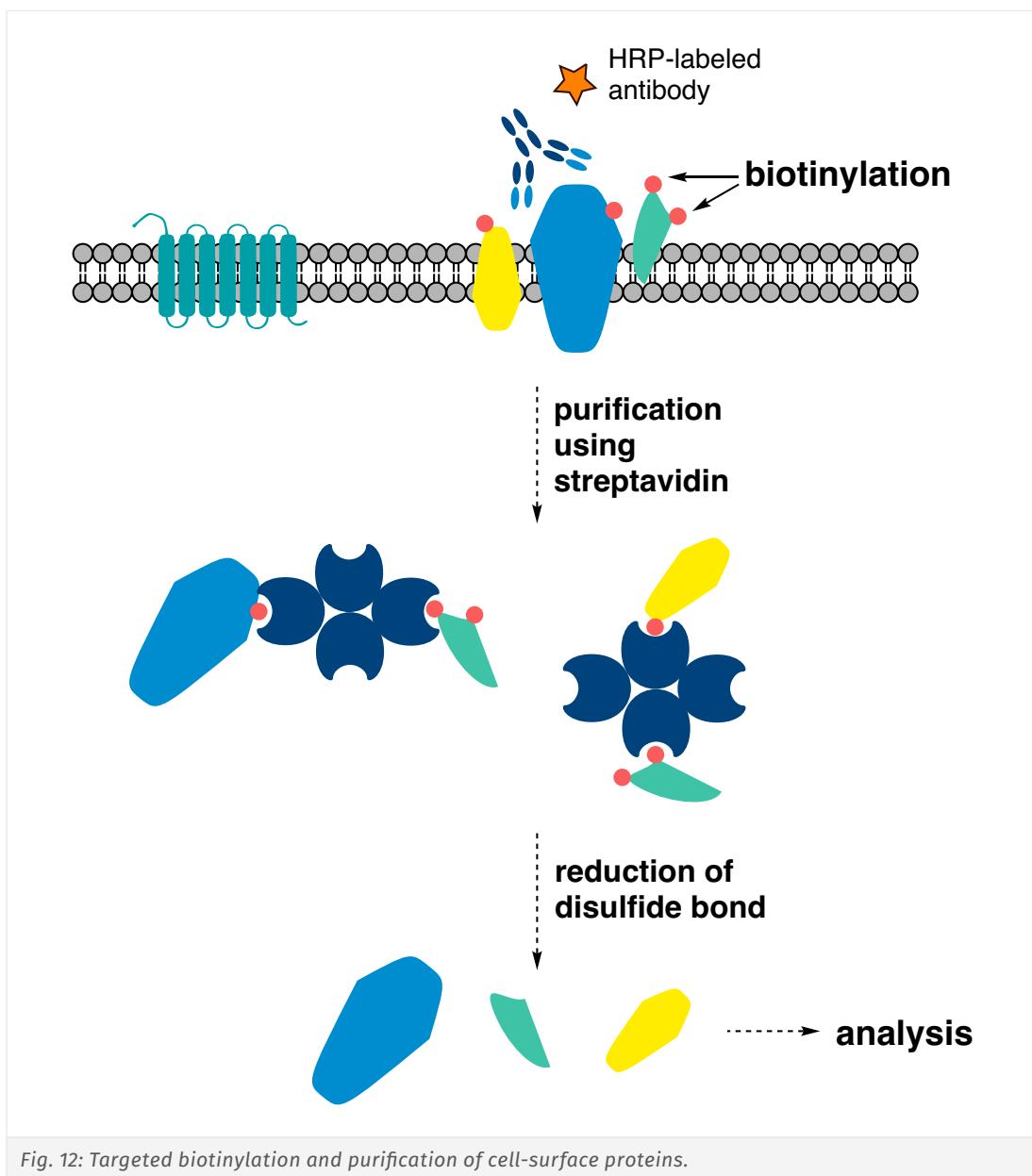
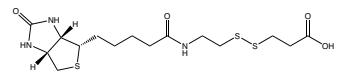


Fig. 12: Targeted biotinylation and purification of cell-surface proteins.

RL-3300 Biotin-SS-COOH

3-((2-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamido)ethyl)disulfanyl)propanoic acid

CAS-No. 104582-29-8
 Formula C₁₅H₂₅N₃O₄S₃
 Mol. weight 407,57 g/mol



Product details



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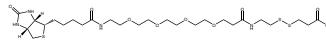
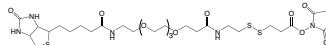
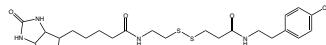
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Biotinylation Reagents

		Product details
PEG8090	Biotin-PEG(4)-SS-COOH	 9,25-dioxo-29-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)-12,15,18,21-tetraoxa-4,5-dithia-8,24-diazanonacosan-1-oic acid CAS-No. 1380166-80-2 Formula C ₂₆ H ₄₆ N ₄ O ₉ S ₃ Mol. weight 654,86 g/mol
PEG1910	Biotin-PEG(4)-S-S-NHS	 1-Biotinamino-15-oxo-3,6,9,12-tetraoxa-19,20-dithia-16-azatricosan-23-oic acid succinimidyl ester CAS-No. 1260247-51-5 Formula C ₃₀ H ₄₉ N ₅ O ₁₁ S ₃ Mol. weight 751,93 g/mol
<p>Biotin-SS-tyramide is a valuable linker for peroxidase-promoted targeted protein biotinylation and protein clusters on cell surfaces. By using peroxidase-tagged antibodies, proteins and protein clusters can be selectively biotinylated, and then isolated using (strept)avidin. The biotin-tag can be subsequently removed using reducing agents (e.g., glutathione). One possible application of biotin-SS-tyramide is the targeted biotinylation of proteins and protein clusters on cell surfaces using HRP-labeled antibodies. Labeled proteins are then purified by using (strept)avidin and released by reductive cleavage of the linker disulfide bond.</p>		
LS-3570	Biotin-SS-Tyramide	 N-(2-((3-(4-hydroxyphenethylamino)-3-oxopropyl)disulfanyl)ethyl)-5-(2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide CAS-No. 678975-20-7 Formula C ₂₅ H ₃₄ N ₄ O ₄ S ₃ Mol. weight 526,74 g/mol
LS-3930	Biotin-PEG(4)-SS-Tyramide	 N-(2-((3-(4-hydroxyphenethylamino)-3-oxopropyl)disulfanyl)ethyl)-1-(5-(2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamido)-3,6,9,12-tetraoxapentadecan-15-amide Formula C ₃₄ H ₅₅ N ₅ O ₉ S ₃ Mol. weight 774,02 g/mol

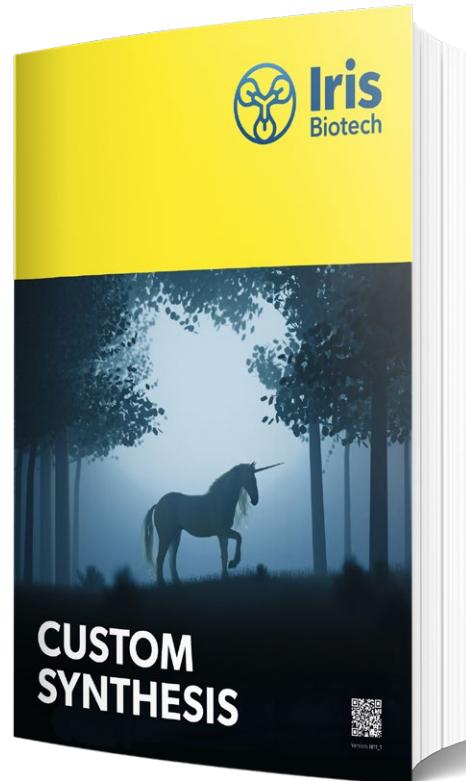
References:

- New insights into the DT40 B cell receptor cluster using a proteomic proximity labeling assay; X. W. Li, J. S. Rees, P. Xue, H. Zhang, S. W. Hamiaux, B. Sanderson, P. E. Funk, R. W. Farndale, K. S. Lilley, S. Perrett, A. P. Jackson; **J Biol Chem** 2014; **289**: 14434-47. ↗ <https://doi.org/10.1074/jbc.M113.529578>
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- Easily reversible desthiobiotin binding to streptavidin, avidin, and other biotin-binding proteins: uses for protein labeling, detection, and isolation; J. D. Hirsch, L. Eslamizar, B. J. Filanowski, N. Malekzadeh, R. P. Haugland, J. M. Beechem, R. P. Haugland; **Anal Biochem** 2002; **308**: 343-57. ↗ [https://doi.org/10.1016/s0003-2697\(02\)00201-4](https://doi.org/10.1016/s0003-2697(02)00201-4)



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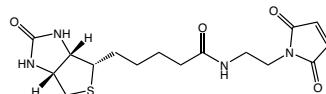
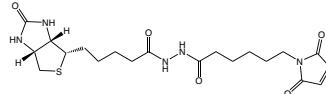
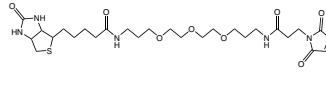
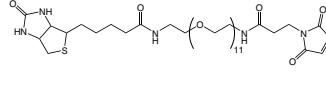
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6. Sulfhydryl-Reactive Biotinylation Reagents

Besides amines, sulfhydryl groups of cysteines are the second most-common targets for biotinylation. Maleimide groups are highly reactive towards sulfhydryl groups at acidic to neutral pH, forming a thioether bond. Alternatively, thiols and pyridyl disulfide-substituted biotin derivatives can be used, resulting in the formation of a cleavable disulfide bond. In the case of using pyridyldisulfides, the biotinylation reaction can visually be monitored by the release of pyridine-2-thione.

Biotin Reagents with Maleimide Function

		Product details
LS-4220	Biotin-mal	 <chem>CC1(C(=O)NCC2=CSC(C2)C1)CC(=O)NCC3=CSC(C3)C(=O)N</chem> N-(2-(2,5-dioxo-2,5-dihydro-1H-pyrrol-1-yl)ethyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide CAS-No. 139554-72-6 Formula C ₁₆ H ₂₂ N ₄ O ₄ S Mol. weight 366,44 g/mol
RL-8420	Biotin-NH-NH-Mal	 <chem>CC1(C(=O)NCC2=CSC(C2)C1)CC(=O)NCC3=CSC(C3)C(=O)NCC4=CSC(C4)C(=O)N</chem> 6-(2,5-dioxo-2,5-dihydro-1H-pyrrol-1-yl)-N'-(5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanoyl)hexanehydrazide CAS-No. 116919-18-7 Formula C ₂₀ H ₂₉ N ₅ O ₅ S Mol. weight 451,54 g/mol
PEG1605	Biotin-PEG(3)-mal	 <chem>CC1(C(=O)NCC2=CSC(C2)C1)CC(=O)NCC3=CSC(C3)C(=O)NCC4=CSC(C4)C(=O)NCC5=CSC(C5)C(=O)N</chem> N-(3-(2-(3-(Biotinamino)propoxy)ethoxy)ethoxy)propyl-3-maleimidylpropanamide CAS-No. 525573-22-2 Formula C ₂₇ H ₄₃ N ₅ O ₉ S Mol. weight 597,72 g/mol
PEG1595	Biotin-PEG(11)-mal	 <chem>CC1(C(=O)NCC2=CSC(C2)C1)CC(=O)NCC3=CSC(C3)C(=O)NCC4=CSC(C4)C(=O)NCC5=CSC(C5)C(=O)NCC6=CSC(C6)C(=O)N</chem> alpha-Biotin-omega-maleimido undeca(ethylene glycol) CAS-No. 1334172-60-9 Formula C ₄₁ H ₇₁ N ₅ O ₁₆ S Mol. weight 922,09 g/mol

Product details

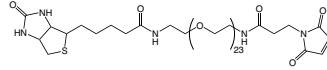
PEG4320 Biotin-PEG(23)-mal

alpha-Biotin-omega-maleimido 23(ethylene glycol)

CAS-No. 1334172-60-9

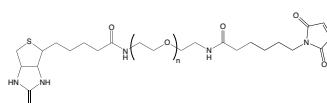
Formula C₆₅H₁₁₉N₅O₂₈S

Mol. weight 1450,72 g/mol


PEG1049 Biotin-PEG-mal (3 kDa)

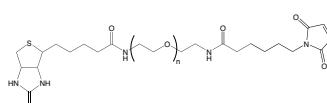
alpha-Biotin-omega-maleimido poly(ethylene glycol)

Mol. weight 3000 Da


PEG1050 Biotin-PEG-mal (5 kDa)

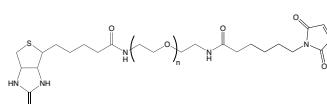
alpha-Biotin-omega-maleimido poly(ethylene glycol)

Mol. weight 5000 Da


PEG1048 Biotin-PEG-mal (10 kDa)

alpha-Biotin-omega-maleimido poly(ethylene glycol)

Mol. weight 10000 Da

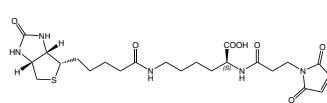

RL-8425 Biocytin-Mal

N2-(3-(2,5-dioxo-2,5-dihydro-1H-pyrrol-1-yl)propanoyl)-
N6-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]
imidazol-4-yl)pentanoyl-L-lysine

CAS-No. 102849-12-7

Formula C₂₃H₃₃N₅O₇S

Mol. weight 523,61 g/mol



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Biotin Reagents with Thiol Function

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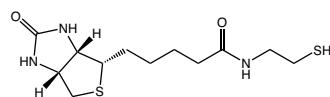
LS-4250 Biotin-Cysteamine

N-(2-mercaptopethyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide

CAS-No. 99938-68-8

Formula C₁₂H₂₁N₃O₂S₂

Mol. weight 303,44 g/mol



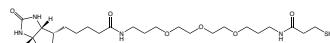
LS-4260 Biotin-PEG(3)-SH

N-(17-mercaptopro-4,7,10-trioxa-14-azadecyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide

CAS-No. 1208242-09-4

Formula C₂₃H₄₂N₄O₆S₂

Mol. weight 534,73 g/mol



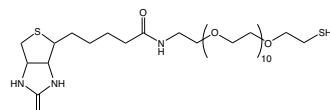
PEG4705 Biotin-PEG(11)-SH

alpha-Biotin-omega-mercaptop undeca(ethylene glycol)

CAS-No. 1650579-23-9

Formula C₃₄H₆₅N₃O₁₃S₂

Mol. weight 788,02 g/mol

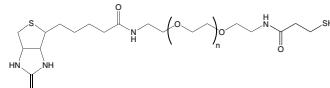


PEG1213 Biotin-PEG-SH (3 kDa)

alpha-Biotin-omega-mercaptop poly(ethylene glycol)

CAS-No. 1030917-83-9

Mol. weight 3000 Da

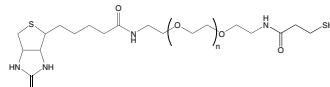


PEG1214 Biotin-PEG-SH (5 kDa)

alpha-Biotin-omega-mercaptop poly(ethylene glycol)

CAS-No. 1030917-83-9

Mol. weight 5000 Da



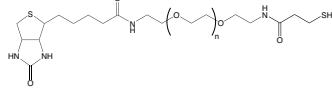
Product details

PEG1212 Biotin-PEG-SH (10 kDa)

alpha-Biotin-omega-mercaptopoly(ethylene glycol)

CAS-No. 1030917-83-9

Mol. weight 10000 Da

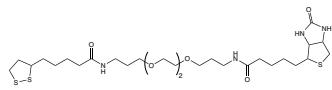

PEG3570 Lipoamide-PEG(3)-biotin

alpha-Lipoamide-omega-biotinyl tri(ethylene glycol)

CAS-No. 1334172-74-5

 Formula C₂₈H₅₀N₄O₆S₃

Mol. weight 634,91 g/mol

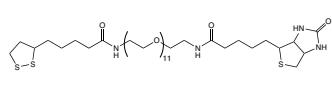

PEG3580 Lipoamide-PEG(11)-biotin

alpha-Lipoamide-omega-biotinyl undeca(ethylene glycol)

CAS-No. 960069-81-2

 Formula C₄₂H₇₈N₄O₁₄S₃

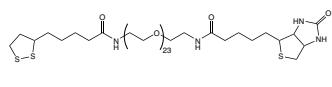
Mol. weight 959,28 g/mol


PEG7530 Lipoamide-PEG(23)-biotin

alpha-Lipoamide-omega-biotinyl 23(ethylene glycol)

 Formula C₆₆H₁₂₆N₄O₂₆S₃

Mol. weight 1487,91 g/mol


Biotin Reagents with Cleavable Disulfide Unit

Product details

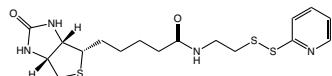
LS-4230 Biotin-SS-Py

5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)-N-(2-(pyridin-2-yl disulfanethyl)ethyl)pentanamide

CAS-No. 112247-65-1

 Formula C₁₇H₂₄N₄O₂S₃

Mol. weight 412,59 g/mol



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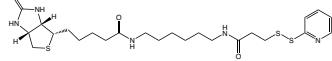
RL-8415 Biotin-Hx-SS-Py

5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)-N-(6-(3-(pyridin-2-yldisulfanyl)propanamido)hexyl)pentanamide

CAS-No. 129179-83-5

Formula C₂₄H₃₇N₅O₃S₃

Mol. weight 539,77 g/mol



LS-4240 Biotin-PEG(3)-SS-Py

N-(15-oxo-17-(pyridin-2-yldisulfanetyl)-4,7,10-trioxa-14-azaheptadecyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide

Formula C₂₈H₄₅N₅O₆S₃

Mol. weight 643,88 g/mol

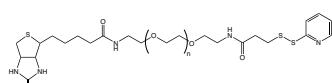


PEG4770 Biotin-PEG-OPSS (3 kDa)

alpha-Biotin-omega-pyridyl-2-disulfid poly(ethylene glycol)

CAS-No. 1776111-43-3

Mol. weight 3000 Da

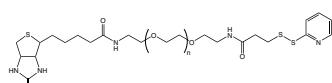


PEG4775 Biotin-PEG-OPSS (5 kDa)

alpha-Biotin-omega-pyridyl-2-disulfid poly(ethylene glycol)

CAS-No. 1776111-43-3

Mol. weight 5000 Da

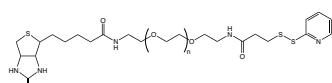


PEG4780 Biotin-PEG-OPSS (10 kDa)

alpha-Biotin-omega-pyridyl-2-disulfid poly(ethylene glycol)

CAS-No. 1776111-43-3

Mol. weight 10000 Da

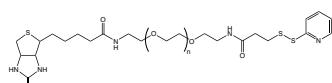


PEG4785 Biotin-PEG-OPSS (20 kDa)

alpha-Biotin-omega-pyridyl-2-disulfid poly(ethylene glycol)

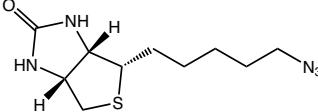
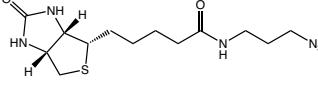
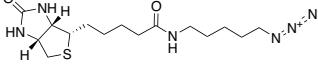
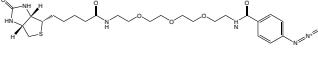
CAS-No. 1776111-43-3

Mol. weight 20000 Da



7. Click-Reactive Biotinylation Reagents

Our portfolio includes azide- as well as alkyne-derivatized biotins suitable for conjugation via click chemistry. As further benefit triazole-linked adducts are shown to be highly resistant to the ubiquitous hydrolytic enzyme biotinidase and to bind (strept)avidin with dissociation constants in the low pM range.

		Product details
LS-4300	DecarboxyBiotin-N₃	 
(3aS,4S,6aR)- 4-(5-Azidopentyl)tetrahydro-1H-Thieno[3,4-d]imidazol-2(3H)-one		
CAS-No.	1260586-88-6	
Formula	C ₁₀ H ₁₇ N ₅ OS	
Mol. weight	255,34 g/mol	
LS-4210	Biotin-N₃	 
N-(3-azidopropyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide		
CAS-No.	908007-17-0	
Formula	C ₁₃ H ₂₂ N ₆ O ₂ S	
Mol. weight	326,42 g/mol	
LS-4660	Biotinyl-DAPe-N₃	 
N-(5-azidopentyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide		
CAS-No.	1349190-76-6	
Formula	C ₁₅ H ₂₆ N ₆ O ₂ S	
Mol. weight	354,47 g/mol	
PEG6795	Biotin-PEG3-Ph(4-N₃)	 
Biotin-PEG3-phenyl azide		
CAS-No.	2088238-77-9	
Formula	C ₂₅ H ₃₇ N ₃ O ₆ S	
Mol. weight	563,67 g/mol	

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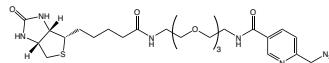
Biotinylation Reagents

Product details

LS-4680 Biotin-PEG(3)-PyAz

6-(azidomethyl)-N-(13-oxo-17-((3aS,4S,6aR)-2-oxo-hexahydro-1H-thieno[3,4-d]imidazol-4-yl)-3,6,9-trioxa-12-azaheptadecyl)nicotinamide

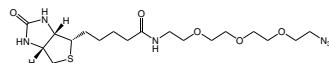
Formula $C_{25}H_{38}N_8O_6S$
Mol. weight 578,69 g/mol



PEG4940 Biotin-PEG(3)-N₃

11-[D(+)-Biotinylamino]-1-azido-3,6,9-trioxaundecane

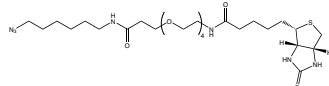
CAS-No. 875770-34-6
Formula $C_{18}H_{32}N_6O_5S$
Mol. weight 444,55 g/mol



PEG7990 Biotin-PEG(4)-N₃

(3aS,4S,6aR)-4-(28-azido-5,21-dioxo-9,12,15,18-te-traoxa-6,22-diazaoctacosyl)tetrahydrono-1H-thieno[3,4-d]imidazol-2(3H)-one

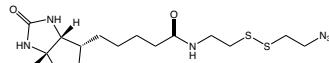
CAS-No. 1006592-62-6
Formula $C_{27}H_{49}N_6O_7S$
Mol. weight 615,79 g/mol



RL-4120 Biotin-SS-N₃

N-(2-((2-azidoethyl)disulfanyl)ethyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide

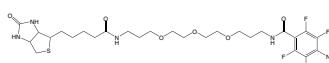
CAS-No. 1620523-64-9
Formula $C_{14}H_{24}N_6O_2S_3$
Mol. weight 404,57 g/mol



PEG2065 Biotin-TEG-ATFBA

Biotin-triethylenglycol-(*p*-azido-tetrafluorobenzamide)

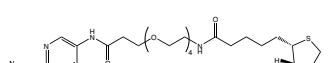
CAS-No. 1264662-85-2
Formula $C_{27}H_{37}F_4N_7O_6S$
Mol. weight 663,68 g/mol



PEG8000 Biotin-PEG(4)-Picolyl-N₃

(3aS,4S,6aR)-4-(1-(6-(azidomethyl)pyridin-3-ylamino)-1,17-dioxo-4,7,10,13-tetraoxa-16-azahenicosan-21-yl)-tetrahydro-1H-thieno[3,4-d]imidazol-2(3H)-one

CAS-No. 2222687-71-8
Formula $C_{27}H_{42}N_8O_7S$
Mol. weight 622,74 g/mol



Product details

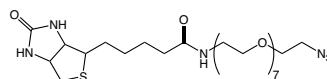
PEG8100 Biotin-PEG(4)-SS-Azide

N-(2-((3-((3-azidopropyl)amino)-3-oxopropyl)disulfanefyl)ethyl)-1-(5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamido)-3,6,9,12-tetraoxapentadecan-15-amide

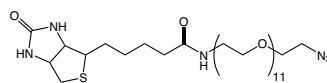
CAS-No. 1260247-52-6
 Formula C₂₉H₅₂N₈O₈S₃
 Mol. weight 736,96 g/mol

**PEG4330 Biotin-PEG(7)-N₃**

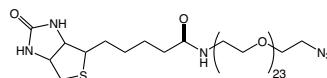
alpha-Biotin-omega-azido hepta(ethylene glycol)
 CAS-No. 1334172-75-6
 Formula C₂₆H₄₈N₆O₉S
 Mol. weight 620,76 g/mol

**PEG4340 Biotin-PEG(11)-N₃**

[2-(2-aminoethoxy)ethoxy]acetic acid *tert*-butyl ester*HCl
 CAS-No. 956494-20-5
 Formula C₃₄H₆₄N₆O₁₃S
 Mol. weight 796,97 g/mol

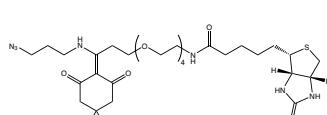
**PEG4350 Biotin-PEG(23)-N₃**

alpha-Biotin-omega-azido 23(ethylene glycol)
 CAS-No. 956494-20-5
 Formula C₅₈H₁₁₂N₆O₂₅S
 Mol. weight 1325,6 g/mol

**PEG7960 Biotin-PEG(4)-Dde-N₃**

N-(19-azido-15-(4,4-dimethyl-2,6-dioxocyclohexylidene)-3,6,9,12-tetraoxa-16-azanonadecyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide

CAS-No. 1802907-93-2
 Formula C₃₂H₅₃N₇O₈S
 Mol. weight 695,87 g/mol



Background

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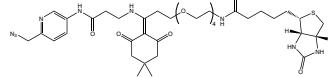
PEG7970 Biotin-PEG(4)-Dde-Picolyl-N₃

N-(6-(azidomethyl)pyridin-3-yl)-15-(4,4-dimethyl-2,6-dioxocyclohexylidene)-1-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamido)-3,6,9,12-tetraoxa-16-azononadecan-19-amide

CAS-No. 2055048-42-3

Formula C₃₈H₅₇N₉O₉S

Mol. weight 815,98 g/mol



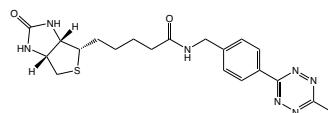
LS-4280 Biotin-MeTz

N-(4-(6-methyl-1,2,4,5-tetrazin-3-yl)benzyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide

CAS-No. 1802883-51-7

Formula C₂₀H₂₅N₇O₂S

Mol. weight 427,53 g/mol



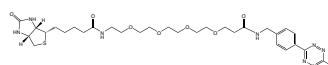
LS-4290 Biotin-PEG(4)-MeTz

N-(4-(6-methyl-1,2,4,5-tetrazin-3-yl)benzyl)-1-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamido)-3,6,9,12-tetraoxapentadecan-15-amide

CAS-No. 1962919-31-8

Formula C₃₁H₄₆N₈O₇S

Mol. weight 674,82 g/mol



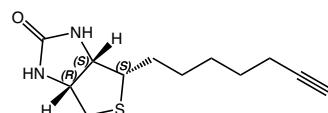
LS-4310 DecarboxyBiotin-Alkyne

(3aS,4S,6aR)-4-(6-heptyn-1-yl)tetrahydro-1H-Thieno[3,4-d]imidazol-2(3H)-one

CAS-No. 887915-53-9

Formula C₁₂H₁₈N₂OS

Mol. weight 238,35 g/mol



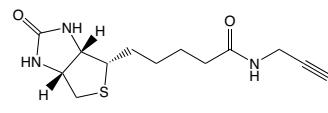
RL-3490 Biotin-Propargylamide

Biotinyl-N-propargylamide

CAS-No. 773888-45-2

Formula C₁₃H₁₉N₃O₂S

Mol. weight 281,37 g/mol



Product details

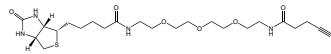
PEG6805 Biotin-PEG3-NH-Pentynoyl

alpha-Biotin-omega-pentynoyl 3(ethylene glycol)

CAS-No. 869082-82-6

Formula C₂₃H₃₈N₄O₆S

Mol. weight 498,64 g/mol

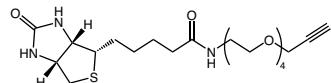
**PEG4950 Biotin-PEG(4)-alkyne**

15-[D(+)-Biotinylamino]-4,7,10,13-tetraoxapenta-dec-1-yne

CAS-No. 1262681-31-1

Formula C₂₁H₃₅N₃O₆S

Mol. weight 457,58 g/mol

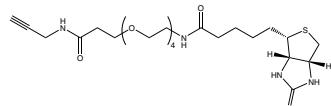
**PEG8010 Biotin-PEG(4)-Alkyne**

(3aS,4S,6aR)-4-(5,21-dioxo-8,11,14,17-tetraoxa-4,20-diazapentacos-1-yn-25-yl)tetrahydro-1H-thieno[3,4-d]imidazol-2(3H)-one

CAS-No. 1006592-45-5

Formula C₂₄H₄₀N₄O₈S

Mol. weight 528,66 g/mol

**PEG8110 Biotin-PEG(4)-SS-Alkyne**

N-(2-((3-oxo-3-(prop-2-ynylamino)propyl)disulfanyl)ethyl)-1-(5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamido)-3,6,9,12-tetraoxapentadecan-15-amide

CAS-No. 1260247-54-8

Formula C₂₉H₄₉N₅O₈S₃

Mol. weight 691,92 g/mol

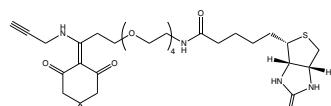
**PEG7980 Biotin-PEG(4)-Dde-Alkyne**

N-(15-(4,4-dimethyl-2,6-dioxocyclohexylidene)-3,6,9,12-tetraoxa-16-azononadec-18-ynyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide

CAS-No. 1802908-00-4

Formula C₃₂H₅₀N₄O₈S

Mol. weight 650,83 g/mol



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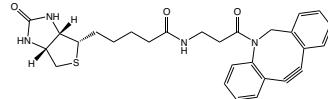
LS-4270 Biotin-DBCO

(3aS,4S,6aR)-N-[3-(11,12-Didehydrodibenz[b,f]azocin-5(6H)-yl)-3-oxopropyl]hexahydro-2-oxo-1H-thieno[3,4-d]imidazole-4-pentanamide

CAS-No. 1418217-95-4

Formula C₂₈H₃₀N₄O₃S

Mol. weight 502,63 g/mol



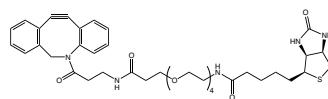
RL-2520 Biotin-PEG(4)-DBCO

Dibenzoazacyclooctyne-tetra(ethylene glycol)-biotin

CAS-No. 1255942-07-4

Formula C₃₉H₅₁N₅O₈S

Mol. weight 749,92 g/mol

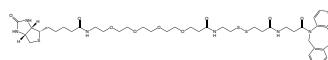


PEG8120 Biotin-PEG(4)-SS-DBCO

N-(2-((3-(3-(azadibenzocyclooctyn-1-yl)-3-oxopropylamino)-3-oxopropyl)disulfanyl)ethyl)-1-(5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamido)-3,6,9,12-tetraoxapentadecan-15-amide

Formula C₄₄H₆₀N₆O₉S₃

Mol. weight 913,18 g/mol



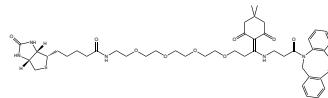
PEG8140 Biotin-PEG(4)-Dde-DBCO

N-(15-(4,4-dimethyl-2,6-dioxocyclohexylidene)-19-oxo-19-(azadibenzocyclooctyn-1-yl)-3,6,9,12-tetraoxa-16-azanonadecyl)-5-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)pentanamide

CAS-No. 1807512-43-1

Formula C₄₇H₆₁N₅O₉S

Mol. weight 872,08 g/mol





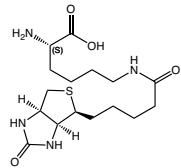
8. Biotinylated Amino Acids

Biocytin is a multi-purpose marker for neuroanatomical studies and the ideal substrate for biotinidase assays. Its strong complex formation with (strept)avidin conjugates opens up a broad range of detection methods. As biocytin is the natural substrate of the enzyme biotinidase, biocytin can be used to measure the biotinidase activity and therefore diagnose biotinidase deficiency. As an intermediate in the metabolism of biotin, biocytin occurs naturally in blood serum and urine.

Product details

LS-3510 Biocytin

N-epsilon-Biotinyl-L-Lysine
CAS-No. 576-19-2
Formula C ₁₆ H ₂₈ N ₄ O ₄ S
Mol. weight 372,48 g/mol



Reference:

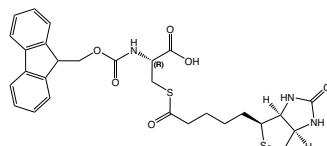
- Improved neuronal tract tracing with stable biocytin-derived neuroimaging agents; A. Mishra, K. Dhingra, A. Schuz, N. K. Logothetis, S. Canals; *ACS Chem Neurosci* 2010; 1: 129-38. <https://doi.org/10.1021/cn900010d>

For the introduction of biotin during peptide synthesis, biotinylated amino acid building blocks based on cysteine, glutamic acid, and lysine are available.

Product details

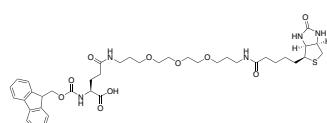
FAA3720 Fmoc-L-Cys(Biotin)-OH

N-alpha-(9-Fluorenylmethyloxycarbonyl)-S-biotinyl-L-cysteine
Formula C ₂₈ H ₃₁ N ₃ O ₆ S ₂
Mol. weight 569,69 g/mol



FAA8430 Fmoc-L-Glu(Biotin-TOTA)-OH

N-alpha-(9-Fluorenylmethyloxycarbonyl)-N-gamma-(N-biotinyl-3-(2-(2-(3-aminopropoxy)ethoxy)ethoxy)propyl)-L-glutamine
CAS-No. 817169-73-6
Formula C ₄₀ H ₅₅ N ₅ O ₁₀ S
Mol. weight 797,96 g/mol



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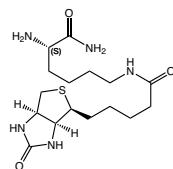
HAA3430 H-L-Lys(Biotin)-NH₂

N-epsilon-biotin-L-lysine amide

CAS-No. 61125-53-9

Formula C₁₆H₂₉N₅O₃S

Mol. weight 371,50 g/mol



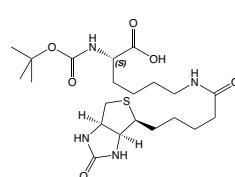
BAA1276 Boc-L-Lys(Biotin)-OH

N-alpha-t-Butyloxycarbonyl-N-epsilon-biotinyl-L-lysine

CAS-No. 62062-43-5

Formula C₂₁H₃₆N₄O₆S

Mol. weight 472,6 g/mol



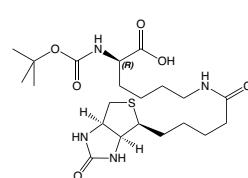
BAA1038 Boc-D-Lys(Biotin)-OH

N-alpha-t-Butyloxycarbonyl-N-epsilon-(Biotin)-D-lysine

CAS-No. 1272755-71-1

Formula C₂₁H₃₆N₄O₆S

Mol. weight 472,61 g/mol



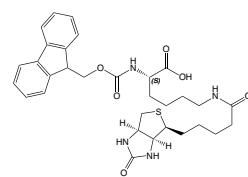
FAA1443 Fmoc-L-Lys(Biotin)-OH

N-alpha-(9-Fluorenylmethyloxycarbonyl)-N-epsilon-biotinyl-L-lysine

CAS-No. 146987-10-2

Formula C₃₁H₃₈N₄O₆S

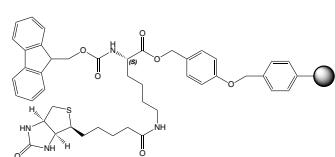
Mol. weight 594,7 g/mol



WAA5154 Fmoc-L-Lys(Biotin)-Wang Resin

Mesh Size 100-200 mesh

DVB 1% DVB



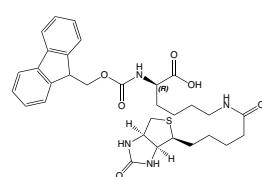
FAA1451 Fmoc-D-Lys(Biotin)-OH

N-alpha-(9-Fluorenylmethyloxycarbonyl)-N-epsilon-biotinyl-D-lysine

CAS-No. 110990-09-5

Formula C₃₁H₃₈N₄O₆S

Mol. weight 594,7 g/mol

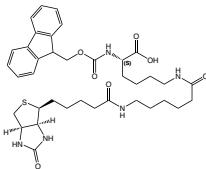


Product details

FAA4670 Fmoc-L-Lys(Biotin-Ahx)-OH

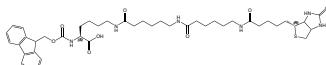
N-alpha-(9-Fluorenylmethyloxycarbonyl)-N-epsilon-
lon-[6-(biotinylamino)hexanoyl]-L-lysine

CAS-No. 160158-05-4
Formula C₃₇H₄₉N₅O₇S
Mol. weight 707,88 g/mol


FAA8765 Fmoc-L-Lys(Biotin-Ahx-Ahx)-OH

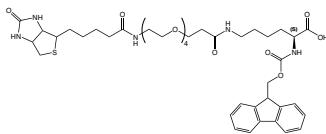
N-alpha-(9-Fluorenylmethyloxycarbonyl)-N-epsilon-
lon-(biotinyl-biscaproyl)-L-lysine

Formula C₄₃H₆₀N₆O₈S
Mol. weight 821,05 g/mol


PEG4440 Fmoc-L-Lys(PEG(4)-Biotin)-OH

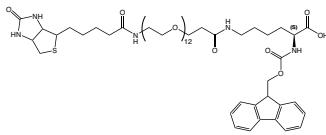
N-alpha-(9-Fluorenylmethyloxycarbonyl)-N-epsi-
lon-[15-(biotinamido)-4,7,10,13-tetraoxa-penta-
decanoyl]-L-lysine

CAS-No. 1334172-64-3
Formula C₄₂H₅₉N₅O₁₁S
Mol. weight 842,01 g/mol


PEG4450 Fmoc-L-Lys(PEG(12)-Biotin)-OH

N-alpha-(9-Fluorenylmethyloxycarbonyl)-N-epsi-
lon-[alpha-Biotin-omega-propionyl dodeca(ethylene
glycol)]-L-lysine

CAS-No. 1334172-65-4
Formula C₅₈H₉₁N₅O₁₉S
Mol. weight 1194,43 g/mol


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Background

Amine-Reactive
Biotinylation Reagents

Carboxy-Reactive
Biotinylation Reagents

Reagents for Various Protein
and RNA Biotinylations

Reversible Biotinylation

Sulphydryl-Reactive
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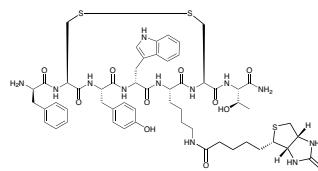
9. Other Biotin Reagents

Product details

LS-1522 TT-232-Biotin diacetate

H-D-Phe-Cys-Tyr-D-Trp-Lys(Biotin)-Cys-Thr-NH₂ (disulfide bridged)

Formula C₅₅H₇₂N₁₂O₁₁S₃ (net)
Mol. weight 1173,43 (net) g/mol



PEG2080 Biotin-TEG-Biotin

N,N'-Bisbiotin-tetra(ethylene glycol)-diamine

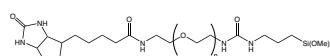
CAS-No. 1263044-47-8
Formula C₃₀H₅₂N₆O₈S₂
Mol. weight 672,9 g/mol



PEG4850 Biotin-PEG-Si(OMe)₃ (3 kDa)

alpha-Biotinyl-omega-trimethoxysilyl poly(ethylene glycol)

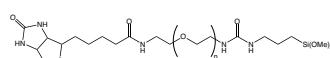
Mol. weight 3000 Da



PEG4855 Biotin-PEG-Si(OMe)₃ (5 kDa)

alpha-Biotinyl-omega-trimethoxysilyl poly(ethylene glycol)

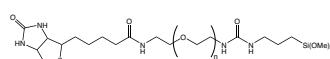
Mol. weight 5000 Da



PEG4860 Biotin-PEG-Si(OMe)₃ (10 kDa)

alpha-Biotinyl-omega-trimethoxysilyl poly(ethylene glycol)

Mol. weight 10000 Da

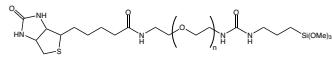


Product details

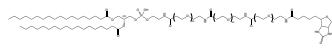
PEG4865 Biotin-PEG-Si(OMe)₃ (20 kDa)

alpha-Biotinyl-omega-trimethoxysilyl poly(ethylene glycol)

Mol. weight 20000 Da

**PEG7125 Biotin-PEG(4)-amido-PEG(24)-amido-PEG(24)-DSPE**

(2R)-3-((hydroxy((4,80,156,172-tetraoxo-176-(2-oxohe-xahydro-1H-thieno[3,4-d]imidazol-4-yl)-7,10,13,16,19,22,25,28,31,34,37,40,43,46,49,52,55,58,61,64,67,70,73,76,83,86,89,92,95,98,101,104,107,110,11

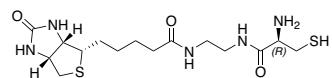
**RL-8645 Biotin-NH-CH₂-CH₂-NH-Cysteine*TFA**

N-Biotinyl-N'-cysteinyl ethylenediamine*TFA

CAS-No. 288144-42-3

Formula C₁₅H₂₇N₅O₃S₂*CF₃CO₂H

Mol. weight 389,53*114,02 g/mol



While nitriles in general may react with sulphydryl groups (e.g., from cysteines) in a rather unspecific way, the advantage of α -cyanopyridines (2-picolinonitrils) is their strong selectivity for 1,2-aminothiols (e.g., provided as N-terminal cysteine or internally as non-canonical amino acid). Thus, this reaction may be used for the synthesis of cyclic peptides or to fuse ligands to peptide chains. It is biorthogonal, biocompatible, catalyst-free, and selective; it proceeds readily in aqueous solutions at physiological pH and ambient temperature. The formation of the resulting 2-thiazoline (alternative name: 2,5-dihydrothiazole) is driven by the release of ammonia.

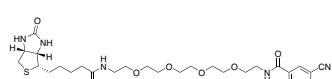
Product details

RL-8635 Biotin-PEG(4)-CINA

2-cyano-N-(16-oxo-20-((3aS,4S,6aR)-2-oxohexahydro-1H-thieno[3,4-d]imidazol-4-yl)-3,6,9,12-tetraoxa-15-azaicosyl)isonicotinamide

Formula C₂₇H₄₀N₆O₅

Mol. weight 592,71 g/mol

**Reference:**

- *The Cyanopyridine-Aminothiol Click Reaction: Expanding Horizons in Chemical Biology; C. Nitsche; SynLett.*
2024; 35: A-E. <https://dx.doi.org/10.1055/a-2214-7612>

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Code of Conduct

As business activity of Iris Biotech GmbH impacts people's lives and health, it must be operated in ethical and correct manner and act with integrity and responsibility. To ensure high ethical standards and fair business practices, Iris Biotech GmbH applies an integrated policy known as its Code of Conduct.

In 2001 Iris Biotech GmbH was founded just at the beginning of the Biotech movement and the first remarkable breakthrough of biotech pharma products. Although the biotech field is rather young compared to other industries we believe on long-term business, a good partnership between our business partners and Iris Biotech GmbH and a good reputation. It is our duty as well as our responsibility to maintain and to extend this over the next generations – based on the principles of an honourable and prudent tradesman which based upon the concept of honourable entrepreneurship.

This Code of Conduct has been developed following the "Voluntary Guidelines for Manufacturers of Fine Chemical Intermediates and Active Ingredients" issued by AIME (Agrochemical & Intermediates Manufacturers in Europe) and the requirements of some of our business associates.

Iris Biotech GmbH commits to hold this Code of Conduct and to include and apply its principles in the management system and the company policies.

Ethics

Iris Biotech GmbH undertakes business in an ethical manner and acts with integrity. All corruption, extortion and embezzlement are prohibited. We do not pay or accept bribes or participate in other illegal inducements in business or government relationships. We conduct our business in compliance with all applicable anti-trust laws. Employees are encouraged to report concerns or illegal activities in the workplace, without threat of reprisal, intimidation or harassment.

Labour

Iris Biotech GmbH is committed to uphold the human rights of workers and to treat them with dignity and respect. Child labour, workplace harassment, discrimination, and harsh and inhumane treatment are prohibited. Iris Biotech GmbH respects the rights of the employees to associate freely, join or not join labour unions, seek representation and join workers' councils. Employees are paid and their working timetable is established according to applicable wage and labour laws. Employees are able to communicate openly with management regarding working conditions without threat of reprisal, intimidation or harassment.

General Policies

Contracts and Secrecy Agreements are binding and the confidential information received is only used for intended purposes. Clear management and organizational structures exist to provide efficient normal working and to address problems quickly. Know-how is protected and intellectual property is respected.

Health and Safety

Iris Biotech GmbH provides a safe and healthy working environment to the employees and protects them from overexposure to chemical and physical hazards. Products are produced, stored and shipped under the guidelines of the relevant chemical and safety legislation. Risks and emergency scenarios are identified and evaluated, and their possible impact is minimized by implementing emergency plans and written procedures. Safety information regarding hazardous materials is available to educate, train and protect workers from hazards. Preventive equipment and facilities maintenance is performed at suitable periods to reduce potential hazards. Employees are regularly trained in health and safety matters and are informed about product properties and risk classification when it is required.

Environment

Iris Biotech GmbH operates in an environmentally responsible and efficient manner, minimizing adverse impacts on the environment. Waste streams are managed to ensure a safe handling, movement, storage, recycling and reuse, before and after being generated. Systems to prevent and mitigate accidental spills and releases to the environment are in place. All required environmental permits and licenses are obtained and their operational and reporting requirements are complied with.

Production and Quality Management

A quality management system following the Good Distribution Practices (GDP rules) of Active Pharmaceutical Ingredients is established covering all the aspects of the worldwide distribution of products. Regular audits are performed to evaluate the efficiency and fulfilling of the quality system. Process controls to provide reproducible product quality are established. There are preventive maintenance procedures to ensure plant reliability and the lowest risk of failure. Staff is trained periodically about GMP and GDP rules. Procedures are established and installations are designed to avoid cross contamination. Batch and analytical records are kept for inspection and audit purposes for suitable periods according guidelines.

Research and Development

Research and development staff education is appropriate to their functional activity and they are trained to develop, optimize and scale-up the processes. Intellectual property is respected and know-how protected. Development of manufacturing processes reflects the principles of the Green Chemistry according to the American Chemical Society Green Chemistry Institute. Animal testing is not used unless alternatives are not scientifically valid or accepted by regulators. If animal testing is carried out, animals are treated so that pain and stress are minimized.

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All orders placed by a buyer are accepted and all contracts are made subject to the terms which shall prevail and be effective notwithstanding any variations or additions contained in any order or other document submitted by the buyer. No modification of these terms shall be binding upon Iris Biotech GmbH unless made in writing by an authorised representative of Iris Biotech GmbH.

Placing of Orders

Every order made by the buyer shall be deemed an offer by the buyer to purchase products from Iris Biotech GmbH and will not be binding on Iris Biotech GmbH until a duly authorised representative of Iris Biotech GmbH has accepted the offer made by the buyer. Iris Biotech GmbH may accept orders from commercial, educational or government organisations, but not from private individuals and Iris Biotech GmbH reserves the right to insist on a written order and/or references from the buyer before proceeding.

There is no minimum order value. At the time of acceptance of an order Iris Biotech GmbH will either arrange prompt despatch from stock or the manufacture/acquisition of material to satisfy the order. In the event of the latter Iris Biotech GmbH will indicate an estimated delivery date. In addition to all its other rights Iris Biotech GmbH reserves the right to refuse the subsequent cancellation of the order if Iris Biotech GmbH expects to deliver the product on or prior to the estimated delivery date. Time shall not be of the essence in respect of delivery of the products. If Iris Biotech GmbH is unable to deliver any products by reason of any circumstances beyond its reasonable control („Force Majeure“) then the period for delivery shall be extended by the time lost due to such Force Majeure. Details of Force Majeure will be forwarded by Iris Biotech GmbH to the buyer as soon as reasonably practicable.

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Prices are subject to change. For the avoidance of doubt, the price advised by Iris Biotech GmbH at the time of the buyer placing the order shall supersede any previous price indications. The buyer must contact the local office of Iris Biotech GmbH before ordering if further information is required. Unless otherwise agreed by the buyer and Iris Biotech GmbH, the price shall be for delivery ex-works. In the event that the buyer requires delivery of the products otherwise than ex-works the buyer should contact the local office of Iris Biotech GmbH in order to detail its requirements. Iris Biotech GmbH shall, at its discretion, arrange the buyer's delivery requirements including, without limitation, transit insurance, the mode of transit (Iris Biotech GmbH reserves the right to vary the mode of transit if any regulations or other relevant considerations so require) and any special packaging requirements (including cylinders). For the avoidance of doubt all costs of delivery and packaging in accordance with the buyer's requests over and above that of delivery in standard packaging ex-works shall be for the buyer's account unless otherwise agreed by both parties. Incoterms 2020 shall apply. Any tax, duty or charge imposed by governmental authority or otherwise and any other applicable taxes, duties or charges shall be for the buyer's account. Iris Biotech GmbH may, on request and where possible, provide quotations for multiple packs or bulk quantities, and non-listed items. Irrespective of the type of request or means of response all quotations must be accepted by the buyer without condition and in writing before an order will be accepted by Iris Biotech GmbH. Unless agreed in writing on different terms, quotations are valid for 30 days from the date thereof. Payment terms are net 30 days from invoice date unless otherwise agreed in writing. Iris Biotech GmbH reserves the right to request advance payment at its discretion. For overseas transactions the buyer shall pay all the banking charges of Iris Biotech GmbH. The buyer shall not

be entitled to withhold or set-off payment for the products for any reason whatsoever. Government/Corporate Visa and MasterCard (and other such credit cards) may be accepted on approved accounts for payment of the products. Personal credit cards are not acceptable. Failure to comply with the terms of payment of Iris Biotech GmbH shall constitute default without reminder. In these circumstances Iris Biotech GmbH may (without prejudice to any other of its rights under these terms) charge interest to accrue on a daily basis at the rate of 2% per month from the date upon which payment falls due to the actual date of payment (such interest shall be paid monthly). If the buyer shall fail to fulfil the payment terms in respect of any invoice of Iris Biotech GmbH Iris Biotech GmbH may demand payment of all outstanding balances from the buyer whether due or not and/or cancel all outstanding orders and/or decline to make further deliveries or provision of services except upon receipt of cash or satisfactory securities. Until payment by the buyer in full of the price and any other monies due to Iris Biotech GmbH in respect of all other products or services supplied or agreed to be supplied by Iris Biotech GmbH to the buyer (including but without limitation any costs of delivery) the property in the products shall remain vested in Iris Biotech GmbH.

Shipping, Packaging and Returns

The buyer shall inspect goods immediately on receipt and inform Iris Biotech GmbH of any shortage or damage within five days. Quality problems must be notified within ten days of receipt. Goods must not be returned without prior written authorisation of Iris Biotech GmbH. Iris Biotech GmbH shall at its sole discretion replace the defective products (or parts thereof) free of charge or refund the price (or proportionate price) to buyer. Opened or damaged containers cannot be returned by the buyer without the written prior agreement of Iris Biotech GmbH. In the case of agreed damaged containers which cannot be so returned, the buyer assumes responsibility for the safe disposal of such containers in accordance with all applicable laws.

Product Quality, Specifications and Technical Information

Products are analysed in the Quality Control laboratories of Iris Biotech GmbH's production partners by methods and procedures which Iris Biotech GmbH considers appropriate. In the event of any dispute concerning reported discrepancies arising from the buyer's analytical results, determined by the buyer's own analytical procedures, Iris Biotech GmbH reserves the right to rely on the results of own analytical methods of Iris Biotech GmbH. Certificates of Analysis or Certificates of Conformity are available at the discretion of Iris Biotech GmbH for bulk orders but not normally for prepack orders. Iris Biotech GmbH reserves the right to make a charge for such certification. Specifications may change and reasonable variation from any value listed should not form the basis of a dispute. Any supply by Iris Biotech GmbH of bespoke or custom product for a buyer shall be to a specification agreed by both parties in writing. Technical information, provided orally, in writing, or by electronic means by or on behalf of Iris Biotech GmbH, including any descriptions, references, illustrations or diagrams in any catalogue or brochure, is provided for guidance purposes only and is subject to change.

Safety

All chemicals should be handled only by competent, suitably trained persons, familiar with laboratory procedures and potential chemical hazards. The burden of safe use of the products of Iris Biotech GmbH vests in the buyer. The buyer assumes all responsibility for warning his employees, and any persons who might reasonably be expected to come into contact with the products, of all risks to person and property in any way connected with the products and for instructing them in their safe handling and use. The buyer also assumes the responsibility for the safe disposal of all products in accordance with all applicable laws.

Uses, Warranties and Liabilities

All products of Iris Biotech GmbH are intended for laboratory research purposes and unless otherwise stated on product labels, in the catalogue and product information sheet of Iris Biotech GmbH or in other literature furnished to the buyer, are not to be used for any other purposes, including but not limited to use as or as components in drugs for human or animal use, medical devices, cosmetics, food additives, household chemicals, agricultural or horticultural products or pesticides. Iris Biotech GmbH offers no warranty regarding the fitness of any product for a particular purpose and shall not be responsible for any loss or damage whatsoever arising there from. No warranty or representation is given by Iris Biotech GmbH that the products do not infringe any letters patent, trademarks, registered designs or other industrial rights. The buyer further warrants to Iris Biotech GmbH that any use of the products in the United States of America shall not result in the products becoming adulterated or misbranded within the meaning of the Federal Food, Drug and Cosmetic Act (or such equivalent legislation in force in the buyer's jurisdiction) and shall not be materials which may not, under sections 404, 505 or 512 of the Act, be introduced into interstate commerce. The buyer acknowledges that, since the products of Iris Biotech GmbH are intended for research purposes, they may not be on the Toxic Substances Control Act 1976 („TSCA“) inventory. The buyer warrants that it shall ensure that the products are approved for use under the TSCA (or such other equivalent legislation in force in the buyer's jurisdiction), if applicable. The buyer shall be responsible for complying with any legislation or regulations governing the use of the products and their importation into the country of destination (for the avoidance of doubt to include, without limitation, the TSCA and all its amendments, all EINECS, ELINCS and NONS regulations). If any licence or consent of any government or other authority shall be required for the acquisition, carriage or use of the products by the buyer the buyer shall obtain the same at its own expense and if necessary produce evidence of the same to Iris Biotech GmbH on demand. Failure to do so shall not entitle the buyer to withhold or delay payment. Any additional expenses or charges incurred by Iris Biotech GmbH resulting from such failure shall be for the buyer's account. Save for death or personal injury caused by negligence of Iris Biotech GmbH, sole obligation of Iris Biotech GmbH and buyer's exclusive remedy with respect to the products proved to the satisfaction of Iris Biotech GmbH to be defective or products incorrectly supplied shall be to accept the return of said products to Iris Biotech GmbH for refund of the actual purchase price paid by the buyer (or proportionate part thereof), or replacement of the defective product (or part thereof) with alternative product. Iris Biotech GmbH shall have no liability to the buyer under or arising directly or indirectly out of or otherwise in connection with the supply of products by Iris Biotech GmbH to the buyer and/or their re-sale or use by the buyer or for any product, process or services of the buyer which in any way comprises the product in contract tort (including negligence or breach of statutory duty) or otherwise for pure economic loss, loss of profit, business, reputation, depletion of brand, contracts, revenues or anticipated savings or for any special indirect or consequential damage or loss of any nature except as may otherwise be expressly provided for in these terms. All implied warranties, terms and representations in respect of the products (whether implied by statute or otherwise) are excluded to the fullest extent permitted by law. The buyer shall indemnify Iris Biotech GmbH for and against any and all losses, damages and expenses, including legal fees and other costs of defending any action, that Iris Biotech GmbH may sustain or incur as a result of any act or omission by the buyer, its officers, agents or employees, its successors or assignees, its customers or all other third parties, whether direct or indirect, in connection with the use of any product. For the avoidance of doubt and in the event that Iris Biotech GmbH supplies bespoke or custom product to the buyer's design or specification, this indemnity shall extend to include any claim by a third party that the manufacture of the product for the buyer or the use of the product by the buyer infringes the intellectual property rights of any third party.

General

Iris Biotech GmbH shall be entitled to assign or sub-contract all or any of its rights and obligations hereunder. The buyer shall not be entitled to assign, transfer, sub-contract or otherwise delegate any of its rights or obligations hereunder. Any delay or forbearance by Iris Biotech GmbH in exercising any right or remedy under these terms shall not constitute a waiver of such right or remedy. If any provision of these terms is held by any competent authority to be invalid or unenforceable in whole or in part the validity of the other provisions of these terms and the remainder of the provision in question shall not be affected. These terms shall be governed by German Law and the German Courts shall have exclusive jurisdiction for the hearing of any dispute between the parties save in relation to enforcement where the jurisdiction of the German Courts shall be non-exclusive.



Get in Contact

**Iris**
Biotech

Iris Biotech GmbH
Adalbert-Zoellner-Str. 1
95615 Marktredwitz
Germany

📞 +49 (0) 9231 97121-0
📠 +49 (0) 9231 97121-99
✉️ info@iris-biotech.de
🌐 www.iris-biotech.de

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